C:\Program Files\Stnexp\Queries\Cuthbertson 10 566 487 clm7.str

| Stack | Sta

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ring nodes :
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chain bonds :
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    15-53
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                  64 - 65
                                 71 - 72
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                                        90-91 91-92 92-93 93-94
                                                                    94-95 95-96
            97-98
                  97-104 97-105 97-106 98-99 99-100 100-101
    96-109
    102-103 102-107 103-108
                              103-115
                                         109-114
                                                  114-115
ring bonds :
    1-2 1-64
               2-3 3-4
                         4-5 4-25
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                         33-68
                                56-57
                                        56-61
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    22-33
    66-68
           69-70
exact/norm bonds :
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    1-2 1-64
              2-3
                    2 - 24
                          3 - 4
                               4-5 4-25 5-6
                                               5-38
                                                               8 – 9
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                  11-51
    10-11
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                                               14 - 29
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                                                             23-74
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                                                      23-63
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                         43-44
                                                      66-68 69-70
           41-42
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                                 43-45
                                        64-66 66-67
                                                                    75-76
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    33-68
            78-79
                  79-80
                         81-82
                                 81-112
                                         82-83
                                                84-85 85-86 87-88 88-89
    77-111
                         94-95
                                 96-109 99-100
                                                 100-101 102-103 102-107
           91-92 93-94
    90-91
    103-115 109-114
exact bonds :
    1-35 3-34
               6-26 7-39
                            9-50 12-28
                                          13-52
                                                15-53 18-31 19-55
                                                                      21-62
    22-23
           35-36 36-37 37-75 39-40 40-41 42-47 44-48 44-49 45-46 52-71
    55-56
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chain nodes :

2.4

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` 64-65 76-110 77-78 80-81 82-113 83-84 86-87 89-90 92-93 95-96 97-98 97-104 97-105 97-106 98-99 101-102 103-108 114-115 normalized bonds : 56-57 56-61 57-58 58-59 59-60 60-61 71-72 71-73

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:CLASS 24:CLASS 25:Atom 26:CLASS 27:CLASS 28:CLASS 29:CLASS 30:Atom 31:CLASS 32:CLASS 33:Atom 34:CLASS 35:CLASS 36:CLASS 37:CLASS 38:CLASS 39:CLASS 33:Atom 34:CLASS 42:CLASS 43:CLASS 44:CLASS 45:CLASS 46:CLASS 40:CLASS 41:CLASS 42:CLASS 43:CLASS 44:CLASS 45:CLASS 46:CLASS 47:CLASS 48:CLASS 49:CLASS 50:CLASS 51:CLASS 52:CLASS 53:CLASS 54:CLASS 55:CLASS 56:Atom 57:Atom 58:Atom 59:Atom 60:Atom 61:Atom 62:CLASS 63:CLASS 64:Atom 65:CLASS 66:Atom 67:CLASS 68:Atom 69:Atom 70:Atom 71:CLASS 72:CLASS 73:CLASS 74:CLASS 75:CLASS 76:CLASS 77:CLASS 78:CLASS 79:CLASS 80:CLASS 81:CLASS 82:CLASS 83:CLASS 84:CLASS 85:CLASS 85:CLASS 87:CLASS 86:CLASS 87:CLASS 86:CLASS 87:CLASS 87:C

Cuthbertson 10 566 487 = Structure of Claim 7, as diagrammed in claim

LOGINID:SSPTAHPY1654 FILE 'HOME' ENTERED AT 10:58:00 ON 22 MAR 2007 => file registry FILE 'REGISTRY' ENTERED AT 10:58:10 ON 22 MAR 2007 Uploading C:\Program Files\Stnexp\Queries\Cuthbertson 10 566 487 clm7.str STRUCTURE UPLOADED => d L1 L1 HAS NO ANSWERS STR *** STRUCTURE DIAGRAM IS NOT AVAILABLE *** Structure attributes must be viewed using STN Express query preparation. => s L1 SSS SAM SAMPLE SEARCH INITIATED 10:59:12 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED -0 TO ITERATE 100.0% PROCESSED 0 ITERATIONS 0 ANSWERS SEARCH TIME: 00.00.01 FULL FILE PROJECTIONS: ONLINE **COMPLETE** BATCH **COMPLETE** PROJECTED ITERATIONS: 0 TO 0 TO PROJECTED ANSWERS: 0 SEA SSS SAM L1 => s L1 SSS FULL FULL SEARCH INITIATED 10:59:23 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED -0 TO ITERATE 0 ANSWERS 100.0% PROCESSED 0 ITERATIONS SEARCH TIME: 00.00.01 O SEA SSS FUL L1 => file stnquide FILE 'STNGUIDE' ENTERED AT 10:59:45 ON 22 MAR 2007

```
C:\Program Files\Stnexp\Queries\Cuthbertson 10 566 487 core7.str
chain nodes :
    23 24 26
                 27
                     28
                         29
                              31
                                  32
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                                               55
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                                                       63
                                                           65
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                                                                    71
                                                                        72
                                                                            73
                                                                                74
ring nodes :
    1 2 3 4
                 5
                    6
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    22 25 30
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                                                               70
chain bonds :
                 3-34 5-38
                              6-26 7-39
                                          8-27
                                                 9-50
                                                       11-51
                                                              12-28 13-52
    1-35 2-24
                                                 22-23
                                                                       36-37
    15-53
          17-54
                  18-31
                          19-55
                                  20-32
                                         21-62
                                                       23-63
                                                               35-36
                                                                              37 - 74
                                  42 - 47
                                                                      45-46
    39-40
           40 - 41
                   41 - 42
                          42 - 43
                                         43 - 44
                                                 43-45
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                                                               44-49
                                                                             52-71
                          71-72
                   66-67
                                  71-73
    55-56
           64-65
ring bonds :
    1-2 1-64
              2-3 3-4
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                                  56-57
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    66-68
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exact/norm bonds :
    1-2 1-64 2-3 2-24
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    10-11 11-12 11-51
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exact bonds :
    1-35 3-34 6-26 7-39 9-50 12-28 13-52
                                                  15-53 18-31 19-55 21-62
           35-36 36-37 37-74 39-40 40-41 42-47 44-48 44-49 45-46 52-71
    22-23
    55-56 64-65
normalized bonds :
                                                71-72
    56-57
           56-61
                   57-58
                          58 - 59
                                  59-60
                                         60-61
                                                        71-73
Match level :
    1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom
    10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:CLASS 24:CLASS 25:Atom
                                                   23:CLASS 24:CLASS 25:Atom
             27:CLASS 28:CLASS 29:CLASS
    26:CLASS
                                             30:Atom 31:CLASS
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33:Atom

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chain nodes :
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   23 24
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ring nodes :
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   22 25 30
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chain bonds :
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ring bonds :
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exact/norm bonds :
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    17 - 54
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    41-42
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                                            66-68
                                                    69-70
exact bonds :
    1-35 3-34
               6-26 7-39 9-50 12-28 13-52
                                               15-53 18-31 19-55 21-62
          35-36 36-37 37-74 39-40 40-41 42-47 44-48 44-49 45-46 52-71
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           64 - 65
normalized bonds :
   56-57
                                59-60
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                                                    71 - 73
           56-61
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Match level :
           2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom
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            11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom
            19:Atom 20:Atom 21:Atom 22:Atom
                                                23:CLASS 24:CLASS 25:Atom
            27:CLASS 28:CLASS 29:CLASS 30:Atom 31:CLASS 32:CLASS
    26:CLASS
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33:Atom

C:\Program Files\Stnexp\Queries\Cuthbertson 10 566 487 core7.str

34:CLASS 35:CLASS 36:CLASS 37:CLASS 38:CLASS 39:CLASS 40:CLASS 41:CLASS 42:CLASS 43:CLASS 44:CLASS 45:CLASS 46:CLASS 47:CLASS 48:CLASS 49:CLASS 50:CLASS 51:CLASS 52:CLASS 53:CLASS 54:CLASS 55:CLASS 56:Atom 57:Atom 58:Atom 59:Atom 60:Atom 61:Atom 62:CLASS 63:CLASS 64:Atom 65:CLASS 66:Atom 67:CLASS 68:Atom 69:Atom 70:Atom 71:CLASS 72:CLASS 73:CLASS 74:CLASS

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 39:CLASS

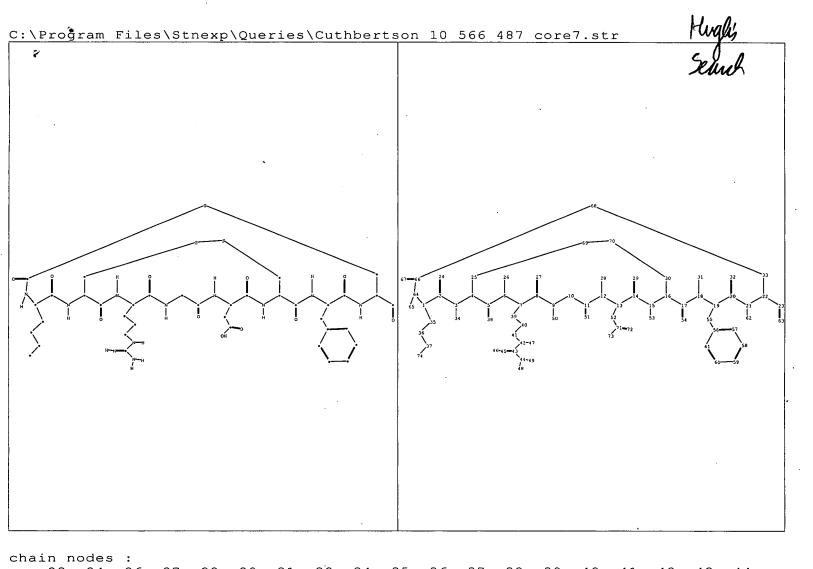
 40:CLASS
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ring nodes :
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chain bonds :
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           64 - 65
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ring bonds :
                          4-5 4-25
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exact/norm bonds :
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                  43 - 44
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                                        66-67
                                                66-68
                                                       69-70
    41 - 42
           42-43
exact bonds :
    1-35 3-34 6-26 7-39 9-50 12-28 13-52
                                                 15-53 18-31 19-55
                                                                        21-62
                  36-37 37-74 39-40 40-41 42-47 44-48 44-49 45-46 52-71
           35-36
    22-23
    55-56
           64 - 65
normalized bonds :
                          58-59
                                 59-60
                                        60-61
                                                71 - 72
    56-57
           56-61
                  57-58
                                                       71-73
Match level :
```

11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:CLASS 24:CLASS 25:Atom

27:CLASS 28:CLASS 29:CLASS 30:Atom 31:CLASS 32:CLASS

8:Atom 9:Atom

23:CLASS 24:CLASS 25:Atom

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom

10:Atom

26:CLASS 33:Atom

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 71:CLASS
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 73:CLASS
 74:CLASS

Cuthbertson 10 566 487 = "core structure" (double-loop) of Claim 7

LOGINID: SSPTAHPY1654 FILE 'HOME' ENTERED AT 10:58:00 ON 22 MAR 2007 => file registry FILE 'REGISTRY' ENTERED AT 10:58:10 ON 22 MAR 2007 Uploading C:\Program Files\Stnexp\Queries\Cuthbertson 10 566 487 clm7.str STRUCTURE UPLOADED => d L1 L1 HAS NO ANSWERS STR *** STRUCTURE DIAGRAM IS NOT AVAILABLE *** Structure attributes must be viewed using STN Express query preparation. => s L1 SSS SAM SAMPLE SEARCH INITIATED 10:59:12 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED -0 TO ITERATE 100.0% PROCESSED 0 ITERATIONS 0 ANSWERS SEARCH TIME: 00.00.01 FULL FILE PROJECTIONS: ONLINE **COMPLETE** BATCH **COMPLETE** PROJECTED ITERATIONS: 0 TO PROJECTED ANSWERS: 0 TO 0 SEA SSS SAM L1 => s L1 SSS FULL FULL SEARCH INITIATED 10:59:23 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED -0 TO ITERATE 100.0% PROCESSED 0 ITERATIONS 0 ANSWERS SEARCH TIME: 00.00.01 0 SEA SSS FUL L1

FILE 'STNGUIDE' ENTERED AT 10:59:45 ON 22 MAR 2007

=> file stnguide

Cuthbertson 10/ 566 487 = STN structure= claim 1/Formula I

Structure-search of Formula I/claim 1; group "W1" is "absent" and not included; group "R2" is present 0-10 repeats, modeled here as (1) repeat or (0 = not present); group R3 is modeled as CH2 (embedded) or as present 1-4 repeats. No stereo-orientations specified.

Searched as SubStructureSearch = SSS sam/ful, LOGINID:SSPTAHPY1654

FILE 'HOME' ENTERED AT 10:15:47 ON 14 MAR 2007

=> file registry

=>

Uploading C:\Program Files\Stnexp\Queries\Cuthbertson 10 566 487 grpI.str

STRUCTURE UPLOADED L1

=> D L1

L1 HAS NO ANSWERS

L1 STR

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

Structure attributes must be viewed using STN Express guery preparation.

=> s L1 SSS SAM

SAMPLE SEARCH INITIATED 10:17:49 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED -0 TO ITERATE

100.0% PROCESSED

0 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

> BATCH **COMPLETE**

PROJECTED ITERATIONS:

0 TO

PROJECTED ANSWERS:

O TO

1.2 0 SEA SSS SAM L1

=> s L1 SSS FULL

FULL SEARCH INITIATED 10:18:04 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 5 TO ITERATE

100.0% PROCESSED

5 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

O SEA SSS FUL L1

=> file stnguide

FILE 'STNGUIDE' ENTERED AT 10:18:15 ON 14 MAR 2007

FILE 'HOME' ENTERED AT 10:51:08 ON 14 MAR 2007

=> file registry

FILE 'REGISTRY' ENTERED AT 10:51:17 ON 14 MAR 2007

Uploading C:\Program Files\Stnexp\Queries\Cuthbertson 10 566 487 sr1b.str

L1 STRUCTURE UPLOADED

=> d L1

L1 HAS NO ANSWERS

L1 STR

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

Structure attributes must be viewed using STN Express query preparation.

=> s L1 sss sam

SAMPLE SEARCH INITIATED 10:52:25 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 0 TO ITERATE

100.0% PROCESSED

0 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH

COMPLETE

PROJECTED ITERATIONS:

0 TO 0

PROJECTED ANSWERS:

0 TO (

L2 0 SEA SSS SAM L1

=> s L1 SSS FULL

FULL SEARCH INITIATED 10:52:35 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED -

3 TO ITERATE

100.0% PROCESSED

3 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

L3 0 SEA SSS FUL L1

FILE 'HOME' ENTERED AT 10:57:17 ON 14 MAR 2007

=> file registry

FILE 'REGISTRY' ENTERED AT 10:57:27 ON 14 MAR 2007

=>

Uploading C:\Program Files\Stnexp\Queries\Cuthbertson 10 566 487 srlc.str

L1 STRUCTURE UPLOADED

=> d L1

L1 HAS NO ANSWERS

L1 STI

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

Structure attributes must be viewed using STN Express query preparation.

=> s L1 SSS SAM

SAMPLE SEARCH INITIATED 10:58:11 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 0 TO ITERATE

100.0% PROCESSED

0 ITERATIONS

O ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 0 TO 0
PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1

=> s L1 SSS FULL FULL SEARCH INITIATED 10:58:19 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 0 TO ITERATE

100.0% PROCESSED 0 ITERATIONS 0 ANSWERS SEARCH TIME: 00.00.01

L3 0 SEA SSS FUL L1

=> file stnguide FILE 'STNGUIDE' ENTERED AT 10:58:33 ON 14 MAR 2007

```
C:\Program Files\Stnexp\Queries\Cuthbertson 10 566 487 grpI.str
chain nodes :
          24
                    27
                         28
                              29
                                        32
                                             34
                                                  35
                                                       36
                                                                 38
                                                                     39
                                                                           40
    23
               26
                                   31
                                                            37
                                                                                41
                                                                                     42
                                                                                         43
                                                                                              44
          46
               47
                         49
                              50
                                   51
                                        52
                                             53
                                                  54
                                                       55
                                                            56
                                                                 57
                                                                          59
     4.5
                    48
                                                                     58
                                                                               60
                                                                                    67
                                                                                              70
                                                                                         68
     72
          77
               78
                    79
                         80
                              81
                                   82
                                        83
                                             8 4
                                                  85
                                                       86
                                                            87
                                                                 88
                                                                     89
                                                                          90
                                                                               91
                                                                                    92
                                                                                         93
                                                                                              94
     95
          96
               97
                    98
                         99
ring nodes :
    1 2 3
                    5
                       6
                           7
                               8
                                   9
                                      10
                                                12
                                                     13
                                                          14
                4
                                           11
                                                               15
                                                                    16
                                                                         17
                                                                              18
                                                                                   19
                                                                                        20
                                                                                             21
     22
         25
               30
                    33
                         61
                              62
                                   63
                                        64
                                             65
                                                  66
                                                     .69
                                                           71
                                                                73
                                                                     74
                                                                          75
                                                                               76
chain bonds
                                          7-41
    1-35
                    3 - 34
                           5-40
                                   6-26
                                                 8-27
                                                         9-52
                                                                 11 - 53
                                                                         12-28
            2-24
                                                                                  13 - 54
                                                                                           14 - 29
    15-58
             17-59
                      18-31
                               19-60
                                        20-32
                                                21-67
                                                         22-23
                                                                          23-77
                                                                  23-68
                                                                                   35-36
                                                                                            36 - 37
                                        43 - 44
     37 - 38
             37-39
                      41 - 42
                               42-43
                                                44 - 45
                                                         44 - 49
                                                                  45-46
                                                                          45 - 47
                                                                                   46-50
                                                                                            46-51
             54 - 55
                      54 - 57
                               55-56
                                                69-70
                                                                  77-78
                                                                          77-79
     47 - 48
                                        60-61
                                                         71 - 72
                                                                                   79-80
                                                                                            80-81
    81-82
             82-83
                      83 - 84
                                                         87-88
                               84 - 85
                                        85-86
                                                86-87
                                                                  88 - 89
                                                                          89 - 90
                                                                                   90-91
                                                                                            90 - 99
             91-96
                      92 - 93
                               93-94
                                        94 - 95
                                                95-97
     91-92
                                                         95 - 98
ring bonds :
     1-2
          1-69
                 2-3
                        3 - 4
                               4-5 4-25
                                            5-6
                                                   6-7
                                                         7-8
                                                               8-9
                                                                     9-10
                                                                             10-11
                                                                                      11-12
                      14-15
                               15-16
                                      16-17
                                                16-30
                                                         17-18
                                                                                   20-21
     12-13
             13 - 14
                                                                  18-19
                                                                          19-20
                                                                                            21-22
     22-33
             25 - 74
                      30-75
                               33-73
                                        61-62
                                                61-66
                                                         62-63
                                                                  63-64
                                                                          64 - 65
                                                                                   65-66
                                                                                            69 - 71
```

71-73

10-11

17-59

33 - 73

75-76

92-93

exact bonds : 1-35

22-23

54 - 55

normalized bonds : 61-62

1-2

exact/norm bonds :

74 - 76

1-69 2-3

11-12

18-19

43 - 44

77-79

93-94

35-36

60-61

61-66

3 - 34

75 - 76

11-53

19-20

44 - 45

80 - 81

95-97

36-37

69-70

62-63

2 - 24

3 - 4

12-13

20-21

45 - 46

81-82

95-98

37-38

77-78

63-64

6-26, 7-41 9-52

4-5

13-14

20-32

45 - 47

83-84

37-39

79-80

64-65

12-28

5-6

13-54

14-15

21 - 22

54 - 57

84 - 85

41-42

82-83

65-66

5-40

14 - 29

22-33

55-56

86-87.

42-43

85-86

15-58

4-25

6-7

15-16

23-68

69-71

87 - 88

44-49

88-89

18-31

7-8

8-9

19-60

16-17

23-77

71-72

89-90

46-50

90-99

8-27

16-30

25 - 74

71 - 73

90-91

46-51

91-92

21-67

9 - 10.

17-18

30 - 75

74 - 76

91-96

47 - 48

94 - 95

Match level :
 1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:CLASS 24:CLASS 25:Atom 26:CLASS 27:CLASS 28:CLASS 29:CLASS 30:Atom 31:CLASS 32:CLASS 33:Atom 34:CLASS 35:CLASS 36:CLASS 37:CLASS 38:CLASS 39:CLASS 40:CLASS 41:CLASS 42:CLASS 43:CLASS 44:CLASS 45:CLASS 46:CLASS 47:CLASS 48:CLASS 49:CLASS 50:CLASS 51:CLASS 52:CLASS 53:CLASS 54:CLASS 55:CLASS 56:CLASS 57:CLASS 58:CLASS 59:CLASS 60:CLASS 61:Atom 62:Atom 63:Atom 64:Atom 65:Atom 66:Atom 67:CLASS 68:CLASS 69:Atom 70:CLASS 71:Atom 72:CLASS 73:Atom 74:Atom 75:Atom 76:Atom 77:CLASS 78:CLASS 80:CLASS 81:CLASS 82:CLASS 83:CLASS 91:CLASS 92:CLASS 93:CLASS 94:CLASS 95:CLASS 97:CLASS 97:CLASS 96:CLASS 97:CLASS 98:CLASS 97:CLASS 97:CLASS 97:CLASS 98:CLASS 97:CLASS 97:CLASS 98:CLASS 98:CLASS

exact/norm bonds : 3 - 44-5 4-25 5-6 5 - 406-7 7-8 8 - 91-2 1-66 2-3 2 - 248-27 9-10 10-11 11-12 11-53 12-13 13 - 1414-15 14 - 2915-16 16-17 16-30 17-18 17-56 18 - 1919-20 20 - 2120 - 3221-22 22-33 23-65 23 - 7325 - 7130 - 7268-70 72-99 33-70 43 - 4444 - 4545-46 45 - 4766-68 68-69 71-99 73-75 79-80 76-77 77-78 85-86 86-87 88-89 80-81 82-83 83-84 87-92 89-90 91 - 9391 - 94exact bonds : 3-34 6-26 7-41 9-52 12-28 1-35 13-54 15-55 18-31 19-57 21 - 6444-49 22-23 35-36 36-37 37 - 3837-39 41-42 42-43 46-50 46-51 47 - 4854-96 57-58 66-67 73 - 7475-76 78-79 81-82 84 - 8586-95 87-88 90-91 normalized bonds : 58-59 58-63 59-60 60-61 61-62 62-63 96-97 96-98

58-63

17-18

59-60

18-19

60-61

19-20

61-62

20-21

62-63

21-22

66-68

16-17

58-59

13-14

25-71

71-99

12-13 22-33

68-70

14-15

30 - 72

72-99

33 - 70

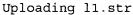
Match level:
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom
10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom
18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:CLASS 24:CLASS 25:Atom
26:CLASS 27:CLASS 28:CLASS 29:CLASS 30:Atom 31:CLASS 32:CLASS
33:Atom 34:CLASS 35:CLASS 36:CLASS 37:CLASS 38:CLASS 39:CLASS
40:CLASS 41:CLASS 42:CLASS 43:CLASS 44:CLASS 45:CLASS 46:CLASS
47:CLASS 48:CLASS 49:CLASS 50:CLASS 51:CLASS 52:CLASS 53:CLASS
54:CLASS 55:CLASS 56:CLASS 57:CLASS 58:Atom 59:Atom 60:Atom 61:Atom
62:Atom 63:Atom 64:CLASS 65:CLASS 66:Atom 67:CLASS 68:Atom 69:CLASS
70:Atom 71:Atom 72:Atom 73:CLASS 74:CLASS 75:CLASS 76:CLASS 77:CLASS
85:CLASS 86:CLASS 87:CLASS 81:CLASS 82:CLASS 83:CLASS 84:CLASS
92:CLASS 93:CLASS 94:CLASS 95:CLASS 96:CLASS 97:CLASS 98:CLASS
99:Atom

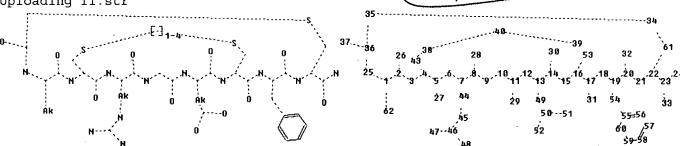
```
chain nodes :
    23 24
            26
                27
                    28
                        29
                            31
                                32
                                    34
                                         35
                                             36
                                                 37
                                                         39
                                                     38
                                                             40
                                                                 41
                                                                     42
                                                                         43
                                                                             44
    4.5
        46
            47
                48
                    49
                        50
                            51
                                52
                                     53
                                         54
                                             55
                                                 56
                                                     57
                                                         64
                                                             65
                                                                 67
                                                                     69
                                                                         73
                                                                             74
    75
ring nodes :
                   6 7
    1 2 3 4
                5
                        8
                            9
                               10
                                   11
                                        12
                                            13
                                                14
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                                                                        20
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                                                                            21
    22 25 30
                33 58
                        59
                            60
                                61
                                    62
                                         63
                                            66
                                                68
                                                     70
                                                         71
                                                             72
                                                                 76
chain bonds :
    1-35 2-24
                3-34 5-40
                            6-26 7-41
                                        8-27
                                               9-52
                                                     11-53
                                                            12-28
                                                                   13-54
                                                                          14-29
    15-55 17-56 18-31
                        19-57
                               20-32
                                       21-64
                                              22-23 23-65 35-36 36-37 37-38
    37-39 41-42
                  42-43
                         43 - 44
                                44 - 45
                                       44 - 49
                                              45-46 45-47
                                                             46-50 46-51
                                                                           47 - 48
                                73-74
                                       73-75
    54 - 73
                  66-67
           57-58
                         68-69
ring bonds :
    1-2 1-66 2-3 3-4
                         4-5 4-25 5-6 6-7
                                              7-8 8-9
                                                        9-10 10-11
                                                                      11 - 12
                         15-16 16-17 16-30
    12-13 13-14
                 14-15
                                              17-18 18-19 19-20 20-21 21-22
    22-33
          25-71
                         33-70
                                58-59
                  30-72
                                       58 - 63
                                               59-60 60-61 61-62
                                                                    62-63 66-68
    68-70
           71-76
                  72-76
exact/norm bonds :
    1-2 1-66 2-3
                    2-24
                         3-4
                               4-5 4-25 5-6
                                               5-40
                                                      6-7 7-8 8-9
                                                                    8-27
                                                                           9-10
    10-11
           11-12
                 11-53
                         12-13
                                13-14
                                       14-15
                                              14-29
                                                      15-16
                                                             16-17
                                                                    16-30
                                                                           17-18
                                       21-22
    17-56
                  19-20
                         20 - 21
                                20-32
                                               22-33
           18-19
                                                      23-65
                                                             25-71
                                                                    30 - 72
                                                                           33 - 70
    43 - 44
           44 - 45
                  45-46
                         45 - 47
                                66-68
                                        68-69
                                               68-70
                                                      71-76
                                                             72-76
exact bonds :
    1-35 3-34
               6-26 7-41 9-52 12-28 13-54
                                                              19-57
                                                15-55 18-31
                                                                      21-64
    22-23
           35-36
                  36-37
                         37-38
                               37-39 41-42 42-43 44-49 46-50 46-51 47-48
           57-58
    54-73
                  66-67
normalized bonds :
           58-63 59-60
                         60-61
                                61-62
                                        62-63
                                              73-74
                                                      73-75
    58-59
Match level :
    1:Atom
            2:Atom 3:Atom
                           4:Atom
                                   5:Atom 6:Atom
                                                     7:Atom
                                                             8:Atom
                                                                     9:Atom
            11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom
                                        22:Atom
                                                  23:CLASS 24:CLASS 25:Atom
    18:Atom
             19:Atom
                      20:Atom
                               21:Atom
```

26:CLASS

C:\Program Files\Stnexp\Queries\Cuthbertson 10 566 487 srlc.str

. 27:CLASS 28:CLASS 29:CLASS 30:Atom 31:CLASS 32:CLASS 33:Atom 34:CLASS 35:CLASS 36:CLASS 37:CLASS 38:CLASS 39:CLASS 40:CLASS 41:CLASS 42:CLASS 43:CLASS 44:CLASS 45:CLASS 46:CLASS 47:CLASS 48:CLASS 49:CLASS 50:CLASS 51:CLASS 52:CLASS 53:CLASS 54:CLASS 55:CLASS 56:CLASS 57:CLASS 58:Atom 59:Atom 60:Atom 61:Atom 62:Atom 63:Atom 64:CLASS 65:CLASS 74:CLASS 75:CLASS 76:Atom





chain nodes :

23 24 26 27 28 29 32 33 37 48 30 31 44 45 46 47 49 50 51 52 54 62

ring nodes :

1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 2.5 34 35 36 38 39 40 43 53 55 56 57 58 59 60 61 chain bonds :

1-62 2-26 5-27 7-44 8-28 11-29 13-49 14-30 17-31 19-54 20-32 22-23 23-

23-33 36-37 44-45 45-46 46-47 46-48 49-50 50-51 50-52 54-55 ring bonds :

1-2 1-25 2-3 3-4 4-5 4-43 5-6 6-7 7-8 8-9 9-10 10-11 11-12 12-13 13-

14-15 15-16 16-17 16-53 17-18 18-19 19-20 20-21 21-22 22-61 25-36 34-35

34-61 35-36

38-40 38-43 39-40 39-53 55-56 55-60 56-57 57-58 58-59 59-60

exact/norm bonds :

1-2 1-25 1-62 2-3 2-26 3-4 4-5 4-43 5-6 5-27 6-7 7-8 7-44 8-9 8-28 9-10 10-11 11-12 11-29 12-13 13-14 13-49 14-15 14-30 15-16 16-17 16-53 17-18 17-31 18-19

19-20 19-54 20-21 20-32 21-22 22-23 22-61 23-24 23-33 25-36 34-35

35-36 36-37

39-53 44-45 45-46 46-47 46-48 49-50 50-51 50-52 54-55 38-40 38-43 39-40 normalized bonds :

55-56 55-60 56-57 57-58 58-59 59-60

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom

11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 19:Atom

20:Atom 21:Atom

22:Atom 23:CLASS 24:CLASS 25:Atom 26:CLASS 27:CLASS 28:CLASS 29:CLASS

30:CLASS 31:CLASS 32:CLASS

33:CLASS 34:Atom 35:Atom 36:Atom 37:CLASS 38:Atom 39:Atom 40:Atom 43:Atom

44:CLASS

45:CLASS 46:CLASS 47:CLASS 48:CLASS 49:CLASS 50:CLASS 51:CLASS 52:CLASS

53:Atom 54:CLASS

55:Atom 56:Atom 57:Atom 58:Atom 59:Atom 60:Atom 61:Atom 62:CLASS

=> d que 121

L15 94 SEA FILE=HCAPLUS ABB=ON PLU=ON ("CUTHBERTSON A"/AU OR

"CUTHBERTSON A C"/AU OR "CUTHBERTSON A F"/AU OR "CUTHBERTSON A F J"/AU OR "CUTHBERTSON A G S"/AU OR "CUTHBERTSON A M"/AU OR "CUTHBERTSON A S"/AU OR "CUTHBERTSON A Z"/AU OR "CUTHBERTSON ALAN"/AU OR "CUTHBERTSON ALAN J S"/AU OR "CUTHBERTSON ALAN S"/AU OR "CUTHBERTSON ALAN S"/AU)

L16	28 SEA FILE=HCAPLUS ABB=ON	PLU=ON ("SOLBAKKEN M"/AU OR "SOLBAKKE
	N MAGNE"/AU)	
L17	19 SEA FILE=HCAPLUS ABB=ON	PLU=ON L15 AND L16
L18	14 SEA FILE=HCAPLUS ABB=ON	PLU=ON L17 AND (AY<2004 OR PY<2004
,	OR PRY<2004)	
L19	32 SEA FILE=HCAPLUS ABB=ON	PLU=ON (L15 OR L16) AND (IMAGING?)
L20	20 SEA FILE=HCAPLUS ABB=ON	PLU=ON L19 AND (AY<2004 OR PY<2004
	OR PRY<2004)	
L21	24 SEA FILE=HCAPLUS ABB=ON	PLU=ON (L20 OR L18)

=> d que 130

L22	437	SEA	CUTHBERTSON A?/AU
L23	74	SEA	SOLBAKKEN M?/AU
L24	49	SEA	L22 AND L23
L25	36	SEA	L24 AND IMAGING?
L26	25	SEA	L25 AND (AY<2004 OR PY<2004 OR PRY<2004)
L27	47	SEA	(L22 OR L23) AND (IMAGING AGENT?)
L28	29	SEA	L27 AND (AY<2004 OR PY<2004 OR PRY<2004)
L29	40	SEA	(L26 OR L28)
L30	29	SEA	L29 AND (IMAGING AGENT?)

=> dup rem 121,130

FILE 'HCAPLUS' ENTERED AT 13:57:33 ON 15 MAR 2007
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FILE 'WPIX' ENTERED AT 13:57:33 ON 15 MAR 2007 COPYRIGHT (C) 2007 THE THOMSON CORPORATION PROCESSING COMPLETED FOR L21 PROCESSING COMPLETED FOR L30

L36

25 DUP REM L21 L30 (28 DUPLICATES REMOVED)

ANSWERS '1-24' FROM FILE HCAPLUS

ANSWER '25' FROM FILE WPIX

=> d ibib abs hitstr retable 136 tot

L36 ANSWER 1 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 1

ACCESSION NUMBER:

2005:472005 HCAPLUS Full-text

DOCUMENT NUMBER:

143:13254

TITLE:

Contrast agent

INVENTOR(S):

Cuthbertson, Alan; Solbakken, Magne

; Lovhaug, Dagfinn

PATENT ASSIGNEE(S):

Amersham Health AS, Norway

SOURCE:

PCT Int. Appl., 30 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KI	ND DATE		DATE			
WO 200504909 WO 200504909	95 A:	2 20050602	WO 2004-NO358				
W: AE,	AG, AL, AM	, AT, AU, AZ,	BA, BB, BG, BR, BW,	BY, BZ, CA, CH,			
CN,	CO, CR, CU	, CZ, DE, DK,	DM, DZ, EC, EE, EG,	ES, FI, GB, GD,			
GE,	GH, GM, HR	, HU, ID, IL,	IN, IS, JP, KE, KG,	KP, KR, KZ, LC,			
LK,	LR, LS, LT	, LU, LV, MA,	MD, MG, MK, MN, MW,	MX, MZ, NA, NI,			
NO,	NZ, OM, PG	, PH, PL, PT,	RO, RU, SC, SD, SE,	SG, SK, SL, SY,			
			UG, US, UZ, VC, VN,				
			NA, SD, SL, SZ, TZ,	· · · ·			
•			TM, AT, BE, BG, CH,				
			IE, IS, IT, LU, MC,				
			CG, CI, CM, GA, GN,				
	SN, TD, TG	,,,	,,,,	2, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3,			
•		2 20060913	EP 2004-808852	20041123 <			
			GB, GR, IT, LI, LU,				
			CZ, EE, HU, PL, SK,				
			US 2005-559880				
PRIORITY APPLN. 1			NO 2003-5228				
			GB 2004-16062				
			WO 2004-NO358				
OTHER SOURCE(S)	MΔ1	PPAT 143.1325		20011123			

OTHER SOURCE(S): MARPAT 143:13254

A contrast agent of formula I: V - L - R (I) where V is an organic group having binding affinity for an angiotensin II receptor site, L is a linear or branched amino acid-comprising biomodifier or linker moiety, and R is a reporter moiety detectable in in vivo imaging of a human or animal body. Contrast agents targeting the AT1 receptor may be suitable for detecting diseases such as congestive heart failure, atherosclerosis, and fibrosis.

L36 ANSWER 2 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 2

2005:471932 HCAPLUS Full-text ACCESSION NUMBER:

DOCUMENT NUMBER: 143:26884

TITLE: Preparation of radiolabeled sulfonamide hydroxamate

matrix metalloproteinase inhibitors as imaging

agents

INVENTOR(S): Cuthbertson, Alan; Solbakken, Magne

; Bjurgert, Emma

PATENT ASSIGNEE(S): Amersham PLC, UK

SOURCE: PCT Int. Appl., 79 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.				KIN	D	DATE			APPLICATION NO.						DATE			
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WO	2005	0490	05		A1		2005	0602	,	WO 2	004-0	GB47	92		2	0041	112 <	
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     AU 2004290950
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                                 20070109
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                                 20070124
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                                                                     20041112 <--
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PRIORITY APPLN. INFO.:
                                             GB 2003-26546
                                                                    20031114 <--
                                             WO 2004-GB4792
                                                                 W
                                                                     20041112
OTHER SOURCE(S):
                         MARPAT 143:26884
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The present invention discloses that <code>imaging</code> agents I [Y1 = H, (CH2)nC(0)Z; n = 1-6; Z = OH, C1-6 alkoxy, C4-10 aryloxy, NR1R2; R1, R2 = independently H, C1-6 alkyl, C3-6 cycloalkyl, C1-6 fluoroalkyl, C4-10 aryl; X1 and X2 form C3-10 cycloalkyl or heterocyclic ring; X3 = H, C1-3 alkyl, C1-3 fluoroalkyl; Y2 = AlpOqA2; p, q = 0-1; Al = C1-10 alkylene, C3-8 cycloalkylene, C1-10 perfluoroalkylene, C6-10 arylene, C2-10 heteroarylene; A2 = H, C1-10 alkyl, C3-8 cycloalkyl, C1-10 perfluoroalkyl, C6-10 aryl, C2-10 heteroaryl] which comprise a specific type of matrix metalloproteinase inhibitors (MMPi's) of the sulfonamide hydroxamate class labeled with an <code>imaging</code> moiety, are useful diagnostic <code>imaging</code> agents for in vivo <code>imaging</code> and diagnosis of the mammalian body. Thus, sulfonamides II (R = iodo) and related compds. were prepared and studies for MMP receptor binding specificity. II and its iodine-123 analog were also studied for biodistribution in a LLC tumor model in vivo, as well as in a ApoE ligation model.

RETABLE

GΙ

Referenced Author	Year	VOL	PG	Referenced Work	Referenced
(RAU)	(RPY)	(RVL)	(RPG)	(RWK)	File
=======================================	+=====-	, -====	, 	, +====================================	+====
Langley, K	2002			,	HCAPLUS

Mobashery, S	2002	US	2002037916 A1	HCAPLUS
Pfizer Products Inc	1999	EF	0895988 A	HCAPLUS
Storey, A	2004	WC	2004069365 A	

L36 ANSWER 3 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 3

ACCESSION NUMBER: 2005:426470 HCAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 142:469186

TITLE: Conjugated angiotensin II analogs as imaging

and therapeutic agents

INVENTOR(S): Cuthbertson, Alan; Indrevoll, Bard; Eriksen,

Morten

PATENT ASSIGNEE(S): Amersham Health A/S, Norway

SOURCE:

PCT Int. Appl., 37 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PA	PATENT NO.			KIND DATE		APPLICATION NO.											
	2005						20050519 20060511							2004			105 <
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CN	1901	1941			Α		2007	0124		CN 2	004-	8003	9758		2	0041	105 <
PRIORIT										NO 2	003-	4952			A 2	0031	106 <
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OTHER SOURCE(S): MARPAT 142:469186

AB The invention comprises pharmaceuticals of formula (I) Z-(L)n-V, wherein V denotes a peptide, L denotes an optional linker, Z denotes a group that optionally can carry an *imaging* moiety M, n denotes 0 or 1. The pharmaceuticals are active as therapeutic agents for the treatment of heart failure, cardiac arrhythmias and diseases where fibrosis is prominent such as COPD, liver fibrosis and atherosclerosis and are also useful as diagnostic agents for the diagnosis of heart failure and diseases were fibrosis is prominent such as COPD, liver fibrosis and atherosclerosis.

L36 ANSWER 4 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 4

ACCESSION NUMBER:

2005:120957 HCAPLUS Full-text

DOCUMENT NUMBER:

142:219561

TITLE:

Preparation of peptide-based compounds as diagnostic

imaging agents

INVENTOR(S):

Cuthbertson, Alan; Solbakken, Magne

PATENT ASSIGNEE(S):

Amersham Health AS, Norway

SOURCE:

PCT Int. Appl., 40 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PATENT NO.								APPLICATION NO.										
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		20061						2006									0060		
		20060				A		2006	0329							_	00602		
PRIORI'	ΤΥ	APPI	NL	LNFO	. :								1781						<
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OTHER : GI	SOU	JRCE	(S):			CASI	REAC	T 14:	2:219	9561	; MAI	RPAT	142	:219	561				

· I

The invention relates to compds. I [R2 is [NH(CH2CH2O)3CH2CH2NHCOCH2CO]0-10NH2; R3 is an alkylene or alkenylene bridge; W1 is absent or a spacer moiety (hetero)hydrocarbyl preferably derived from glutaric and/or succinic acid and/or a polyethylene glycol-based unit and/or a unit [NH(CH2CH2O)3CH2CH2NHCOCH2CO]n; Z1 is an antineoplastic agent, a chelating agent or a reporter moiety] and their use as targeting vectors that bind to receptors associated with angiogenesis. Compds. I may thus be used for diagnosis or therapy of various diseases. Thus, compound I [R2 is NH2, R3 is CH2, Z1-W1 is FCH2CH2SCH2CONH(CH2CH2O)5CH2CH2NHCOCH2COH2CONH] was prepared via the solid-phase method and showed Ki = 7 nmol in an $\alpha v\beta 3$ integrin receptor binding assay.

RETABLE

Referenced Author (RAU)	Year (RPY)	VOL (RVL)	PG (RPG)	Referenced Work (RWK)	Referenced File
Bonasera, T	2002	 	 	WO 02062819 A	HCAPLUS
Harris, T	1996	6	1741	BIOORGANIC & MEDICIN	HCAPLUS
Indrevoll, B	2001	İ	ĺ	WO 0177145 A	HCAPLUS
Indrevoll, B	2003	ĺ	ĺ	WO 03006491 A	HCAPLUS
Lister-James, J	1999			US 5888474 A	HCAPLUS
Pearson, D	1996	39	1372	JOURNAL OF MEDICINAL	HCAPLUS
Srinivasan, A	2002			WO 0220610 A	HCAPLUS

L36 ANSWER 5 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 5

ACCESSION NUMBER: 2005:34777 HCAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 142:130349

TITLE: Fluorescein-labeled peptides

INVENTOR(S): Cuthbertson, Alan; Indrevoll, Bard;

Solbakken, Magne

PATENT ASSIGNEE(S): Amersham Health A/S, Norway

SOURCE: PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PA	PATENT NO.					KIND DATE				APPL:	ICAT:	ION I	NO.	DATE			
WO	2005	0031	66		A1	A1 20050113			1	WO 2	004-1	NO20	В		2	0040	707 <
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									Ţ	WO 2	004-1	NO20	В	1	W 2	0040	707

AB The invention relates to new peptide-based compds. and their use in diagnostic optical *imaging*. More specifically the invention relates to the use of such peptide-based compds. as targeting vectors that bind to receptors associated with angiogenesis. The compds. are labeled with fluorescein and may be used as contrast agents in optical *imaging* in diagnosis of angiogenesis-related diseases.

RETABLE

Referenced Author (RAU)	Year	VOL (RVL)	PG (RPG)	Referenced Work (RWK)	Referenced File
Cuthbertson, A de Groot, F	2002	 1	 901	WO 0226776 A MOLECULAR CANCER THE	HCAPLUS
Hellebust, H Indrevoll, B	2002	į	į	US 2002102217 A1	HCAPLUS
Katada, J	1997	 272	 7720	THE JOURNAL OF BIOLO	!
Riecke, B Univ Leipzig	2001 1999	33 	307 	HORMONE AND METABOLI DE 19808591 A	HCAPLUS HCAPLUS

L36 ANSWER 6 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 6

ACCESSION NUMBER:

2004:610037 HCAPLUS Full-text

DOCUMENT NUMBER:

141:145687

TITLE:

Contrast agents for imaging angiotensin II

receptors

INVENTOR (S):

Solbakken, Magne; Engell, Torgrim;

Wadsworth, Harry John; Archer, Colin M.

PATENT ASSIGNEE(S):

Amersham Health As, Norway

SOURCE:

PCT Int. Appl., 31 pp.

booken.

CODEN: PIXXD2

DOCUMENT TYPE: LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA'	TENT	NT NO. KIND DATE					TE APPLICATION NO.							DATE			
																	•
WO	2004	0625	68		A2	200	40729	1	WO 2	004-1	NO2			2	0040	109	<
WO	2004	0625	68		A3	200	40930										
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EP	1581	262			A2	200	51005		EP 2	004-	7011	36		2	0040	109	<
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CN	1723	043			Α	200	60118	(CN 2	004-	8000	1954		2	0040	109	< ~ -
JP	2006	5175	39		T	200	60727	,	JP 2	006-	5007	37		2	0040	109	<
IN	2005	DN02	527		Α	200	61229		IN 2	005-1	DN25	27		2	0050	511	<
US	2007	0201	76		A1	200	70125	1	US 2	006-	5419	49	•	2	0060	507	<
PRIORIT	Y APP	LN.	INFO	. :]	NO 2	003-3	115		Ž	A 2	0030	109	<
								1	WO 2	004-1	NO2		į	W 2	0040	109	

OTHER SOURCE(S): MARPAT 141:145687

AB The present invention relates to contrast agents in which the targeting vector binds to angiotensin II receptors. The targeting vector, a receptor antagonist such as losartan, valsartan, candesartan or eprosartan, is conjugated via a spacer or linker to a moiety detectable in in vivo imaging procedures. The imaging moiety is a chelated radionuclide such as 99mTc. The imaging moiety may also consist of paramagnetic or fluorescent metal ions or other detectable species.

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10566487
L36 ANSWER 7 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 7
ACCESSION NUMBER:
                        2003:58113 HCAPLUS Full-text
DOCUMENT NUMBER:
                        138:122862
                        Preparation of peptide-based compounds as diagnostic
TITLE:
                        imaging agents
INVENTOR(S):
                        Cuthbertson, Alan; Indrevoll, Bard;
                        Solbakken, Magne
PATENT ASSIGNEE(S):
                        Amersham Health AS, Norway
SOURCE:
                        PCT Int. Appl., 56 pp.
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
LANGUAGE:
                        English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
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     WO 2003006491
                         A2
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PRIORITY APPLN. INFO.:
                                           GB 2001-16815
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OTHER SOURCE(S): MARPAT 138:122862
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R^{1}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2}_\frac{1}{2
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AB Peptide-based compds. I [G represents glycine; D represents aspartic acid; R1 = (CH2)1-10 or (CH2)1-10C6H4, n = 1 or 2; X1 represents an amino acid residue which possesses a functional side chain such as an acid or amine; X2, X4 represent an amino acid residue capable of forming a disulfide bond; X3

represents arginine, N-methylarginine or an arginine mimetic; X5 represents a hydrophobic amino acid or derivative; X6 represents a thiol-containing amino acid residue; X7 is absent or represents a biomodifier moiety; Z1 represents an antineoplastic agent, a chelating agent or a reporter moiety; W1 is absent or represents a spacer moiety] or pharmaceutically-acceptable salts were prepared for use as diagnostic *imaging* agents or as therapeutic agents which comprise targeting vectors which bind to integrin receptors. Thus, cyclo[CH2CO-Lys(cPn216- glutaryl)-Cys2-Arg-Gly-Asp-Cys6-Phe-Cys]-NH2 disulfide (Cys2-6) [cPn216 is technetium chelate residue (HON:CMeCMe2CH2CH2)2CHCH2CH2NH] was prepared via the solid-phase method.

L36 ANSWER 8 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 8

ACCESSION NUMBER:

2003:242904 HCAPLUS Full-text

DOCUMENT NUMBER:

139:328246

TITLE:

Amphiphilic lipopeptide microparticles as contrast

agents for medical ultrasound imaging

AUTHOR (S):

Cuthbertson, Alan; Tornes, Audun;

Solbakken, Magne; Moen, Ove; Eriksen, Morten

CORPORATE SOURCE:

Dep. of Exploratory Res., Amersham Health AS, Oslo,

Norway

SOURCE:

Macromolecular Bioscience (2003), 3(1),

11-17

CODEN: MBAIBU; ISSN: 1616-5187 Wiley-VCH Verlag GmbH & Co. KGAA

DOCUMENT TYPE:

Journal

LANGUAGE:

PUBLISHER:

English

AB In this study the authors investigated the utility of complementary amphiphilic lipopeptides as a new membrane formulation suitable for the preparation of gas-filled microbubbles. Through primarily ion pairing and hydrophobic interactions we rationalized that the stacking of synthetic lipopeptides into the surface of microbubbles would make bubble suspensions useful as ultrasound contrast agents. By mixing charged lipopeptides in propylene glycol/glycerol solns. in the presence of a perfluorocarbon gas followed by vigorous shaking, microbubble suspensions were formed in good yield with a size distribution spanning the range 1-7+10-6 m. The microbubbles were studied in an in vivo model and provided *imaging* efficacy comparable with conventional ultrasound contrast agents.

RETABLE

Km rrap pp					
Referenced Author	Year	VOL	PG	Referenced Work	Referenced
(RAU)	(RPY)	(RVL)	(RPG)	(RWK)	File
=======================================	+====-	+====	+=====	+=============	+=======
Anon	1997		1	Ultrasound Contrast	
Atherton, E	1989			Solid Phase Synthesi	
Blomley, M	2001	322	1222	BMJ	MEDLINE
Bodansky, M	1994			The Practice of Pept	
Church, C	1995	97	1510	J Acoust Soc Am	
Cristiansen, C	1994	19	307	Biotechnol Appl Bioc	
Epstein, P	1950	18	1505	J Chem Phys	HCAPLUS
Fritz, T	1997	32	735	Invest Radiol	HCAPLUS
Fritzsch, T	1988	23	302	Invest Radiol	
Hoff, L	2001			Acoustic Characteriz	
Hoff, L	1996	2	1441	IEEE Ultrasonics Sym	
Hogg, J	1987	67	1249	Physiol Rev	HCAPLUS
Kamp, O	2001	22	1485	Eur Heart J	MEDLINE
Meltzer, R	1980	6	263	Ultrasound Med Biol	MEDLINE
Powers, J	2000	2	15	2nd Symp on Ultrasou	
Unger, E	1998	81	Sympo	Am J Cardiol	
Yang, E	1971	4	283	J Biomech	

10566487 L36 ANSWER 9 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 9 ACCESSION NUMBER: 2002:695821 HCAPLUS Full-text DOCUMENT NUMBER: 137:237702 Improved peptide-chelate conjugates TITLE: INVENTOR(S): Cuthbertson, Alan; Mendizabal, Marivi; Dixon, Mark; Storey, Anthony Eamon PATENT ASSIGNEE(S): Amersham PLC, UK SOURCE: PCT Int. Appl., 50 pp. CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. WO 2002-GB857 WO 2002070018 A2 20020912 20020301 <--WO 2002070018 **A**3 20021205 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, DE GM, HR, HU, ID, IL, IN, IS, JP, LS, LT, LU, LV, MA, MD, MG, MK, PL, PT, RO, RU, SD, SE, SG, SI, UA, UG, US, UZ, VN, YU, ZA, ZM, RW: GH, GM, KE, LS, MW, MZ, SD, SL, CY, DE, DK, ES, FI, FR, GB, GR, BF, BJ, CF, CG, CI, CM, GA, GN, CA 2439579 A1 20020912 EP 1368064 A2 20031210 R: AT, BE, CH, DE, DK, ES, FR, GB, IE, SI, LT, LV, FI, RO, MK, CY, JP 2004524323 ${f T}$ 20040812 US 2006222593 A1 20061005 US 2004-469801 20041210 <--PRIORITY APPLN. INFO.: GB 2001-5224 A 20010302 <--W 20020301 <--WO 2002-GB857 OTHER SOURCE(S): MARPAT 137:237702

AB A peptide-chelate conjugate with affinity for the ST receptor is disclosed, wherein the chelate is tetradentate. The peptide-chelate conjugate of the invention may be labeled with a radiometal to provide a metal complex. A radiopharmaceutical composition comprising the metal complex is provided, which is suitable for the diagnostic *imaging* of colorectal cancer. Also provided for in the invention is a kit for the preparation of the radiopharmaceutical preparation

L36 ANSWER 10 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 10

ACCESSION NUMBER:

2002:256296 HCAPLUS Full-text

DOCUMENT NUMBER:

136:263481

TITLE:

SOURCE:

Preparation of peptide-based compounds as diagnostic

imaging agents

INVENTOR(S):

Cuthbertson, Alan

PATENT ASSIGNEE(S):

Nycomed Imaging AS, Norway; Amersham Health AS

PCT Int. Appl., 39 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.

KIND DATE

APPLICATION NO.

DATE

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WO 2002026776
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                                  20020404
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     EP 1358206
                           A2
                                  20031105
         R: AT, BE, CH, DE, DK, ES, FR, G
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                                  20030324
     US 2003176639
                                  20030918
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PRIORITY APPLN. INFO.:
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                                                                   P
                                              US 2001-259919P
                                                                       20010105 <--
                                              WO 2001-NO390
                                                                   W 20010925 <--
OTHER SOURCE(S):
                        MARPAT 136:263481
GI
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 $[(x^8)_{q}-x^7]_{p}-R^1-x^1-x^2-x^3-G-D-x^4-x^5-x^6-[x^7-(x^8)_{q}]_{p'}$

Peptide-based compds. I [G represents glycine; D represents aspartic acid; p, AΒ p' = 0, 1 and p + p' = 1; when p = 0 then R1 is (CH2)nCO or (CH2)nC6H4CO, where n = 1-5, when p = 1 then R1 is or one or more bridge-forming amino acids; X1 = a bond or 1-5 amino acids, an amino acid derivatized with a carbohydrate moiety, an amino acid functionalized with a spacer or linker and/or a chelate binding or capable of binding a reporter suitable for in vivo imaging; X2, X4 are cysteine, homocysteine or other amino acids capable of forming a cyclizing bond such as aspartic acid and lysine; X3 is arginine, Nmethylarginine, or an arginine mimetic; X5 is a hydrophobic amino acid; X6 is an amino acid capable of forming a cyclizing bond; X7 is a bond or 1-10 amino acids or a spacer or linker, optionally allowing for labeling with multiple chelates as defined by X8, and optionally comprising one or more ethylene glycol units or any other spacer component; X8 is a chelate binding to, or capable of binding, a metal radionuclide or any other reporter suitable for in vivo imaging, NH2 or is absent; q is 0-8; one of the bridges (between R1 and X2 or between X4 and X6) comprises a disulfide bond] were prepared for use in therapeutically effective treatments and as diagnostic imaging agents. More specifically, the invention relates to the use of such peptide-based compds. used as targeting vectors that bind to receptors associated with angiogenesis, in particular the $\alpha v\beta 3$ integrin receptor. Synthesis and conjugation of peptide vector H-Ala-Cys-Asp-Cys-Arg-Gly-Asp-Cys-Phe-Cys-Gly-OH with disulfide bonds connecting Cys-2 and Cys-4 and Cys-8 and Cys-10 and technetium chelatesuccinic acid intermediate [HON:CMeCMe2NHCH2CH2]2NCH2CH2NHCOCH2CH2 CO2H are described.

L36 ANSWER 11 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 11

ACCESSION NUMBER: 2002:386498 HCAPLUS Full-text

DOCUMENT NUMBER: 138:52011

TITLE: In vivo *imaging* of human colon cancer

xenografts in immunodeficient mice using a guanylyl

cyclase C-specific ligand

AUTHOR(S): Wolfe, Henry R.; Mendizabal, Marivi; Lleong, Elinor;

Cuthbertson, Alan; Desai, Vinay; Pullan,

Shirley; Fujii, Dennis K.; Morrison, Matthew; Pither,

Richard; Waldman, Scott A.

CORPORATE SOURCE: Research and Development Department, Targeted

Diagnostics and Therapeutics, Inc., West Chester, PA,

19380, USA

SOURCE: Journal of Nuclear Medicine (2002), 43(3),

392-399

CODEN: JNMEAQ; ISSN: 0161-5505

PUBLISHER: Society of Nuclear Medicine

DOCUMENT TYPE: Journal LANGUAGE: English

AΒ Guanylyl cyclase C (GC-C) is a transmembrane receptor expressed by human intestinal cells and primary and metastatic colorectal adenocarcinomas but not by extraintestinal tissues or tumors. The Escherichia coli heat-stable enterotoxin analog, STa (5-18), is a 14-amino acid peptide that selectively binds to the extracellular domain of GC-C with subnanomolar affinity. This study examined the utility of a radiolabeled conjugate of STa (5-18) to selectively target and image extraintestinal human colon cancer xenografts in vivo in nude mice. The STa conjugate, ethoxyethylmercaptoacetamidoadipoylglycylglycine-STa (5-18) (NC100586), was synthesized and labeled with 99mTc to produce 99mTc-NC100586. This compound was i.v. administered to nude mice bearing human colon cancer xenografts, and specific targeting was evaluated by biodistribution and gamma camera imaging. nude mice, biodistribution and scintigraphic imaging analyses showed selective uptake of 99mTc-NC100586 into human colon cancer xenografts that express GC-C but not into normal tissues that do not express GC-C. Similarly, 99mTc-NC100586 injected i.v. into CD-1 nude mice with human colon cancer hepatic metastases selectively accumulated in those metastases, and .apprx.5-mm foci of tumor cells were visualized after ex vivo imaging of excised livers. Accumulation of 99mTc-NC100586 in human colon cancer xenografts reflected binding to GC-C because 99mTc-NC100588, an inactive analog that does not bind to GC-C, did not selectively accumulate in cancer xenografts compared with normal tissues. Also, coadministration of excess unlabeled STa (5-18) prevented accumulation of 99mTc-NC100586 in human colon cancer xenografts. Furthermore, 99mTc-NC100586 did not selectively accumulate in Lewis lung tumor xenografts, which do not express GC-C. This study showed that i.v. administered STa (5-18) selectively recognizes and binds to GC-C expressed by human colon cancer cells in vivo. Also shown was the ability to exploit this selective interaction to target imaging agents to extraintestinal human colon tumors in nude mice. These results suggest the utility of STa and GC-C for the development of novel targeted imaging and therapeutic agents with high specificity for metastatic colorectal tumors in humans.

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Referenced Author (RAU)	Year (RPY)	, ,	, ,	Referenced Work (RWK)	Referenced File
Almenoff, J	1993	8	865	Mol Microbiol	HCAPLUS
Barany, G	1993	85	106	Int J Pept Protein R	
Blend, M	1998	16	431	Cancer Invest	MEDLINE
Bustin, S	1999	79	1813	Br J Cancer	MEDLINE
。Cagir, B	1999	131	805	Ann Intern Med	HCAPLUS

Carrithers, S	1996	39	171	Dis Colon Rectum	MEDLINE
Carrithers, S	1994	107	1653	Gastroenterology	MEDLINE
Carrithers, S	1996	93	14827	Proc Natl Acad Sci	HCAPLUS
Cohen, M	1988	94	367	Gastroenterology	MEDLINE
Cuthbertson, A	2000	41	3661	Tetrahedron Lett	HCAPLUS
Deshmane, S	1997	36	12921	Biochemistry	HCAPLUS
Erlichmann, C	1994	13	A562	Proc Am Soc Clin Onc	!
Field, M	1978	75	2800	Proc Natl Acad Sci	HCAPLUS
Gariepy, J	1987	84	8907	!	HCAPLUS
Gelmann, A	2000	1	737	Expert Opin Pharmaco	•
Gold, R	1988	36	1	Drugs	
Granowska, M	1993	20	691	Eur J Nucl Med	İ
Greenlee, R	2000	50	7	CA Cancer J Clin	MEDLINE
Guarino, A	1987	32	1017	Dig Dis Sci	HCAPLUS
Guerrant, R	1980	142	220	J Infect Dis	HCAPLUS
Hughes, J	1978	271	755	Nature	HCAPLUS
Hugues, M	1991	30	10738	Biochemistry	HCAPLUS
Ikemura, H	1984	57	2550	Bull Chem Soc	HCAPLUS
Kasina, S	1998	9	108	Bioconjug Chem	HCAPLUS
Krause, B	1998	24	72	Eur J Nucl Med	
Liu, S	1997	8	621	Bioconjug Chem	HCAPLUS
Mayer, R	1992	67	454	Proc Royal Soc Med	
Murakami, H	1980	77	3464	Proc Nat Acad Sci	HCAPLUS
Ohlsson, B	1993	159	275	Eur J Surg	MEDLINE
Raderer, M	1998	39	1570	J Nucl Med	HCAPLUS
Rao, M	1980	632	35	Biochim Biophys Acta	HCAPLUS
Shapiro, S	1992	75	1252	Cancer	
Smart, C	1992	75	1246	Cancer	
Stahl, W	1995	38	2799	J Med Chem	HCAPLUS
Taddei-Peters, W	1992	52	2603	Cancer Res	HCAPLUS
Urbanski, R	1995	1245	29	Biochim Biophys Acta	HCAPLUS
Valk, P	1999	134	503	Arch Surg	MEDLINE
Waldman, S	1998	41	310	Dis Colon Rectum	MEDLINE
Weinberg, D	2000	11	1	Semin Gastrointest D	:
Yamasaki, S	1990	63	2063	Bull Chem Soc	HCAPLUS

L36 ANSWER 12 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 12

ACCESSION NUMBER:

2001:763033 HCAPLUS Full-text

DOCUMENT NUMBER:

135:318716

TITLE:

Preparation of peptide-based compounds as diagnostic

imaging agents

INVENTOR(S):

Cuthbertson, Alan; Indrevoll, Bard

PATENT ASSIGNEE(S):

Nycomed Imaging AS, Norway PCT Int. Appl., 63 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND DA	APPLI APPLI	CATION NO.	DATE
WO 2001077145	A2 20	0011018 WO 20	001-NO146	20010406 <
WO 2001077145	A3 20	0020510		
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YU, ZA, 2	M			

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                         A2
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                                20050629
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     PT 1272507
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PRIORITY APPLN. INFO.:
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                                            GB 2000-25070
                                            WO 2001-NO146
                                                               W 20010406 <--
OTHER SOURCE(S):
                         MARPAT 135:318716
GI
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Peptide-based compds. I [G represents glycine; D represents aspartic acid; R1 AR = (CH2)n or (CH2)nC6H4 (n = 1-10), m = 1 or 2; p = 1-10; X1 represents a bondor 1-5 amino acid residues which can independently be derivatized with a functional side chain suitable for modifying pharmacokinetics or blood clearance rates and can bind a reporter (R) moiety suitable for in vivo imaging via a linker (L) moiety, a chelating agent or an L moiety attached to a chelating agent; X2, X4 represent an amino acid residue capable of forming a disulfide bond; X3 represents arginine, N-methylarginine or an arginine mimetic; X5 represents a hydrophobic amino acid or derivative; X6 represents a thiol-containing amino acid residue; X7 represents an L moiety or 1-10 amino acid residues, optionally as part of an L moiety, with the properties of X1; X7 is absent; X8 represents an R moiety or NH2 or is absent] or pharmaceutically acceptable salts were prepared for use as diagnostic imaging agents or as therapeutic agents which comprise a targeting vector which binds to receptors associated with integrin receptors. Thus, [Cys2-6]cyclo[CH2CONH-Asp-Cys-Arg-Gly-Asp- Cys-Phe-Cys]-Gly-NH(CH2CH2O)2CH2CH2NHCO(CH2)3CH(NHCOCH2SCHMeOEt)CO-Gly-Gly-OH (VIa) was prepared by solid-phase peptide coupling, cyclization, deprotection, and conjugation with N3S-adipate chelator active ester. Compd.VIa was labeled with technetium.

L36 ANSWER 13 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 13

ACCESSION NUMBER:

2001:519335 HCAPLUS Full-text

DOCUMENT NUMBER:

135:111977

TITLE:

Diagnostic/therapeutic agents having

phospholipid-based microbubbles coupled to one or more

vectors

INVENTOR(S):

Klaveness, Jo; Rongved, Pal; Hogset, Anders;
Tolleshaug, Helge; Naevestad, Anne; Hellebust,

Halldis; Hoff, Lars; Cuthbertson, Alan;

Lovhaug, Dagfinn; Solbakken, Magne

PATENT ASSIGNEE(S): Nycomed Imaging As, Norway

10

U.S., 89 pp., Cont.-in-part of U.S. Ser. No. 958,993. SOURCE:

CODEN: USXXAM

DOCUMENT TYPE:

LANGUAGE:

Patent

FAMILY ACC. NUM. COUNT:

English

PATENT INFORMATION)N:	
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PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
US 6261537	B1	20010717			19971029 <
CN 1234742 .	A	19991110			19971028 <
HU 9904595	A2	20000428			19971028 <
US 6331289	B1	20011218			19971028 <
AT 318618	T	20060315			19971028 <
EP 1442751	A1	20040804			19980424 <
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IE, FI, CY					
ES 2224379	Т3	20050301			19980424 <
KR 2000052829	Α	20000825	KR 1999-703658		19990427 <
US 20.02102215	A1	20020801	US 2001-765614		20010122 <
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US 6680047	B2	20040120			
CN 1440816	Α	20030910	CN 2002-160420		20021230 <
US 2004141922	.A1	20040722	US 2003-722075		20031126 <
US 2005002865	A1	20050106	US 2003-734730		20031215 <
US 2007036722	A1	20070215	US 2006-498651		20060803 <
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			US 1997-49263P	P	19970607 <
			US 1997-49266P	P	19970607 <
			US 1997-959206	Α	19971028 <
			US 1997-960054	A1	19971029 <
			EP 1998-917461	А3	19980424 <
			US 2001-765614	В1	20010122 <
			US 2001-925715		20010810 <
			US 2003-722075		20031126 <
AB Targetable diagnos	tic and	or theren	entically active agent		a ultragound

AΒ Targetable diagnostic and/or therapeutically active agents, e.g. ultrasound contrast agents, having reporters comprise gas-filled microbubbles stabilized by monolayers of film-forming surfactants, the reporter being coupled or linked to at least one vector. The gas is air, nitrogen, oxygen, carbon dioxide, hydrogen, an inert gas, a sulfur fluoride, selenium hexafluoride, a low mol. weight hydrocarbon, a ketone, an ester, a halogenated low mol. weight hydrocarbon or their mixts. The film-forming surfactant material is one or more phospholipids selected from the group consisting of phosphatidylserines,

phosphatidylqlycerols, phosphatidylinositols, phosphatidic acids and cardiolipins. A therapeutic agent is an antineoplastic agent, blood product, biol. response modifier, antifungal agent, hormone or hormone analog, vitamin, enzyme, antiallergic agent, tissue factor inhibitor, platelet inhibitor, coagulation protein target inhibitor, fibrin formation inhibitor, fibrinolysis promoter, antiangiogenic, circulatory drug, metabolic potentiator, antitubercular, antiviral, vasodilator, antibiotic, anti-inflammatory, antiprotozoal, antirheumatic, narcotic, opiate, cardiac qlycoside, neuromuscular blocker, sedative, local anesthetic, general anesthetic or genetic material. For example, an endothelial cell adhesion of phosphatidylserine-encapsulated perfluorobutane microbubbles coated with polylysine was higher than adhesion of uncoated microbubbles. Also, a thrombus was detected by ultrasound in patients with suspected venous thrombosis using i.v. phosphatidylserine-encapsulated microbubbles. The microbubbles contained inactivated human thrombin-succinyl-PEG 3400distearoylphosphatidylethanol amine incorporated into the encapsulating membrane.

RETABLE

RETABLE				•	
Referenced Author	Year	AOL	PG	Referenced Work	Referenced
(RAU)	(RPY)	(RVL)	(RPG)	(RWK)	File
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Anon	1991	1		WO 9115244	HCAPLUS
Anon	1993			WO 9320802	HCAPLUS
Anon	1994			CA 2145505	HCAPLUS
Anon .	1994			WO 9407539	HCAPLUS
Anon	1994			WO 9428873	HCAPLUS
Anon	1994			WO 9428874	HCAPLUS
Anon	1995			WO 9503356	HCAPLUS
Anon	1995			WO 9503357	HCAPLUS
Anon	1995			WO 9507072	HCAPLUS
Anon	1995			WO 9515118	HCAPLUS
Anon	1996			EP 0727225	HCAPLUS
Anon	1996			US 08640464	
Anon	1996	·		WO 9639149	HCAPLUS
Anon	1996			WO 9640277	HCAPLUS
Anon	1996			WO 9640285	HCAPLUS
Anon	1996			WO 9641617	HCAPLUS
Anon	1997	,		WO 9723855	
Anon	1997			WO 9733474	HCAPLUS
Anon	1997	Ì	ļ	WO 9741898	HCAPLUS
Anon	1998	!		DE 19626530	HCAPLUS
Anon	1998			WO 9800172	HCAPLUS
Anon	1998			WO 9804293	HCAPLUS
Anon	1998			WO 9819705	HCAPLUS
Anon	1998	1		WO 9820856	HCAPLUS
Anon	1998	1		WO 9842384	HCAPLUS
Elmaleh	1998			US 5716594	HCAPLUS
Friden	1992			US 5154924	HCAPLUS
Grinstaff	1996			US 5505932	
Grinstaff	1997			US 5650156	HCAPLUS
Grinstaff	1997	Ì		US 5665383	HCAPLUS
Klibanov	1997	38	113	Acta Radiologica	İ
Lanza	1997	1		US 5612057	HCAPLUS
Lanza	1997	1		US 5690907	HCAPLUS
Lanza	1998			US 5780010	1
Matsueda	1990			US 4927916	HCAPLUS
McEver	1993			US 5198424	HCAPLUS
Muzykantov	1994	35	1358	J Nuclear Medicine	MEDLINE
Porter	1998			US 5849727	HCAPLUS
Schneider	1997	1		US 5643553	

Tait	1997	US 5632983	HCAPLUS
Thomas, F	1999	Microparticle Prepar	
Torchilin	1996	US 5534241	İ
Tournier	1999	US 5910300	HCAPLUS
Unger	1997	US 5656211	HCAPLUS
Unger	1998	US 5733572	HCAPLUS
Unger	1998	US 5846517	HCAPLUS
Woodle	1991	US 5013556	HCAPLUS
Woodle	1994	US 5356633	HCAPLUS
Worthington Biochemical	1972	Worthington Enzyme M	

L36 ANSWER 14 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 14

ACCESSION NUMBER:

2000:253997 HCAPLUS Full-text

DOCUMENT NUMBER:

132:284295

TITLE:

Contrast agents

INVENTOR(S):

Klaveness, Jo; Naevestad, Anne; Cuthbertson,

Alan; Solbakken, Magne

PATENT ASSIGNEE(S):

Nycomed Imaging As, Norway

SOURCE:

U.S., 31 pp., Cont.-in-part of PCT 9818497.

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 10

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
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US 6051207	Α	20000418	US 1999-300436		19990428 <
US 6331289	B1	20011218	US 1997-959206		19971028 <
ES 2206689	T 3	20040516	ES 1997-910517		19971028 <
US 2002102217	A1	20020801	US 2001-925715		20010810 <
US 6680047	B2	20040120			
US 2005002865	A1	20050106	US 2003-734730		20031215 <
PRIORITY APPLN. INFO.:			GB 1996-22364	Α	19961028 <
			GB 1996-22365	Α	19961028 <
			GB 1996-22366	Α	19961028 <
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			GB 1996-22369	Α	19961028 <
			GB 1997-699	Α	19970115 <
			GB 1997-2195	Α	19970204 <
			GB 1997-9088	Α	19970502 <
			US 1997-48054P	P	19970530 <
			WO 1997-GB2657	A2	19971028 <
			GB 1997-8265	Α	19970424 <
			GB 1997-11837	Α	19970606 <
			GB 1997-11839	Α	19970606 <
			US 1997-49264P	P	19970606 <
			US 1997-49263P	P	19970607 <
			US 1997-49266P	P	19970607 <
			US 1997-959206	Α	19971028 <
			US 2001-925715	A1	20010810 <
001100 0011000 /0\					

OTHER SOURCE(S): MARPAT 132:284295

AB The invention provides a composition containing compds. with a nonpeptide organic group having binding affinity for an endothelin receptor site, a linker moiety or a bond, and a moiety detectable in in vivo *imaging* of a human or animal body. This composition of matter may be used to image diseases and disorders, particularly of the cardiovascular system. A compound was prepared from lysine and 27-0-3-[2-(3-carboxyacryloylamino)-5-

hydroxyphenyl]acryloyloxymycerone and the resulting compound treated with DTPA dianhydride to give a compound which was chelated with Gd or 99mTc.

RETABLE

Referenced Author	Year	VOL	PG	Referenced Work	Referenced
(RAU)	(RPY)	(RVL)	(RPG)	(RWK)	File
=======================================	+=====	+=====	+=====	+======================================	+========
Anon	1991	}		WO 9115244	HCAPLUS
Anon	1994			EP 0606683	HCAPLUS
Anon	1994			CA 2156620	HCAPLUS
Anon	1994			DE 4311023	HCAPLUS
Anon	1994	}	1	AU B-5314694	
Anon	1996	[HCAPLUS
Anon	1996	}		DE 19503644	
Anon	1996	1		CA 2211364	HCAPLUS
Anon	1997			DE 19536781	HCAPLUS
Anon	1997	[DE 19536785	HCAPLUS
Chan	1994	201	228	Biochem Biophys Res	MEDLINE
Us National Library Of	1		l .	Database Medline	

L36 ANSWER 15 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 15

ACCESSION NUMBER:

1999:708651 HCAPLUS Full-text

DOCUMENT NUMBER:

131:319900

TITLE:

Diagnostic/therapeutic agents comprising

membrane-forming amphiphilic lipopeptide-stabilized

gas microbubbles

INVENTOR(S):

Cuthbertson, Alan; Solbakken, Magne

; Wolfe, Henry Raphael

PATENT ASSIGNEE(S):

Marsden, John Christopher, UK; Nycomed Imaging A/S

SOURCE:

PCT Int. Appl., 59 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

	PATENT NO.				KIND DATE			APPLICATION NO.						DATE					
		9955 9955				A2					WO 1	999-	GB12	47		1:	9990	422	<
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												LS,							
												SE,							
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									508	,	JP 2	000-	5455	79		1:	9990	422	<
									116	i	AU 1	999-	3618	7		1:	9990	423	<
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	NO	2000	0.053	82		Α	:	2000	1218	1	NO 2	000-	5382			2	0001	026	<
PRIC	RITY	APP	LN.	INFO	. :					(GB 1	998-	9084		7	A 1	9980	428	<
										1	WO 1	999-0	GB12	47	1	N .1	9990	422	<

AB Novel membrane-forming amphiphilic lipopeptides comprise one or more peptide moieties containing 2-50 aminoacyl residues and one or more hydrocarbon chains containing 5-50 carbon atoms. Such lipopeptides may be used in the formation of stabilized gas microbubble dispersions suitable for use as diagnostic and/or therapeutic agents, for example as ultrasound contrast agents. Perfluorobutane-containing microbubbles were prepared that used N-[3-(2-aminoethanamido)-5-[2-(n-hexadecyl)octadecanamido]benzoyl]gly cine (preparation given) as the membrane-forming agent.

L36 ANSWER 16 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 16

ACCESSION NUMBER:

1999:690991 HCAPLUS Full-text

DOCUMENT NUMBER:

131:308623

TITLE:

Ultrasound imaging contrast agents,

particularly for perfusion in the myocardium

INVENTOR(S):

Eriksen, Morten; Tolleshaug, Helge; Skurtveit, Roald;

Cuthbertson, Alan; Ostensen, Jonny; Frigstad,

Sigmund; Rongved, Pal

PATENT ASSIGNEE(S):

Marsden, John Christopher, UK; Nycomed Imaging AS

SOURCE:

PCT Int. Appl., 80 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA	PATENT NO.					D DATE			DATE						
WO	9953	963				1999	1028						1	9990	 422 <
	W:	ΑE,	AL,	AM,	AT,	AU, AZ,	BA,	BB, BG	, BR,	BY,	CA,	CH,	CN,	CU,	CZ,
		DE,	DK,	EE,	ES,	FI, GB,	GD,	GE, GH	, GM,	HR,	HU,	ID,	IL,	IN,	IS,
		JP,	ΚE,	KG,	KP,	KR, KZ,	LC,	LK, LR	, LS,	LT,	LU,	LV,	MD,	MG,	MK,
		MN,	MW,	MX,	NO,	NZ, PL,	PT,	RO, RU	, SD,	SE,	SG,	SI,	SK,	SL,	TJ,
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		CI,	CM,	GA,	GN,	GW, ML,	MR,	NE, SN	, TD,	TG					
CA	2329	175			A 1	1999	1028	CA	1999-	2329	175		1	9990	422 <
AU	AU 9936172				Α	1999	1108	AU	1999-	3617	2		1	9990	422 <
BR	BR 9909822				Α	2000	1219	BR	BR 1999-9822					9990	422 <
EP	1073	473			A 1	2001	0207	EP	1999-	9181	33		1	9990	422 <
	R:	AT,	BE,	CH,	DE,	DK, ES,	FR,	GB, GR	, IT,	LI,	LU,	NL,	SE,	MC,	PT,
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HU	2001	0287	8		A2	2002	0328	HU	2001-	2878			1	9990	422 <
JP	2002	5122	06		T	2002	0423	JP	2000-	5443	66		1	99904	422 <
IN	2000	00MM	503		Α	2005	0715	IN	2000-	MN50	3		2	0001	011 <
ZA	2000	0057	89		Α	2001	0730	ZA	2000-	5789			2	0001	018 <
NO	2000	0052	50		Α	2000	1218	NO	2000-	5250			2	0001	019 <
US	2004	1464	62		A1	2004	0729	US	2003-	7171	96		2	0031	119 <
PRIORIT	Y APP	LN.	INFO	. :				GB	1998-	8599		Ž	A 1	99804	422 <
								US	1998-	8488	OΡ	1	P 1	9980!	508 <
								WO	1999-	GB12:	21	1	W 1	9990	422 <
									2000-						023 <
AB U]	ltrasc	nic	visu	aliz	atio	n of a	subje	ct, pa:	cticul	larly	of	perf	usio	n in	the

AB Ultrasonic visualization of a subject, particularly of perfusion in the myocardium and other tissues, is performed using novel gas-containing contrast agent prepns. which promote controllable and temporary growth of the gas phase in vivo following administration and can therefore act as deposited perfusion tracers. The prepns. comprise an injectable aqueous medium comprising dispersed gas and an injectable oil-in-water emulsion in which the oil phase comprises a diffusible component capable of diffusion in vivo into the

dispersed gas to promote temporary growth thereof, such that material present at the surfaces of the dispersed gas phase and material present at the surfaces of the dispersed oil phase have affinity for each other, e.g. as a result of having opposite charges. In cardiac perfusion <code>imaging</code> the prepns. may advantageously be coadministered with vasodilator drugs such as adenosine in order to enhance the differences between return signal intensity from normal and hypoperfused myocardial tissue resp. A neg.-charged perfluorobutane gas dispersion and a pos.-charged perfluorodimethylcyclobutane emulsion were simultaneously injected i.v. into a dog. The resulting myocardial contrast effect was far more intense than that observed when the dispersion and emulsion were both neg.-charged. The contrast lasted for 20 min.

RETABLE

Referenced Author (RAU)	Year (RPY)	VOL (RVL)	PG (RPG)	Referenced Work (RWK)	Referenced File
=======================================	+====-	+====-	+=====	+=== ====== ========	+=====================================
Caminati, G	1998	327-3	37	THIN SOLID FILMS	HCAPLUS
Cockbain Julian	1998			WO 9818497 A	HCAPLUS
Epand, R	1997	43	15	BIOPHYSICAL STUDIES	HCAPLUS
Imarx Pharmaceutical Co	1997			WO 9740858 A	HCAPLUS
Iskandrian, A	1994	1	94	JOURNAL OF MUCLEAR C	MEDLINE
Maletinska, L	1996	79	2023	HELV CHIM ACTA	HCAPLUS
Maletinska, L	1997	43	3271	J MED CHEM	
Marsden John Christophe	1998			WO 9818495 A	HCAPLUS
Marsden John Christophe	1998			WO 9818498 A	HCAPLUS
Ono, S	1992	597	293	JOURNAL OF CHROMATOG	
Porter, T	1997	96	L-401	CIRCULATION	
Shehata, A	1997	80	716	AMERICAN JOURNAL OF	HCAPLUS
Unger, E	1993			US 5228446 A	HCAPLUS
Varadarajan, R	1996			US 5580575 A	HCAPLUS

L36 ANSWER 17 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 17

ACCESSION NUMBER:

1998:300865 HCAPLUS Full-text

DOCUMENT NUMBER:

129:4871

TITLE:

Preparation of targetable diagnostic and therapeutic gas-containing or gas-generating ultrasound contrast

agents

INVENTOR(S):

Klaveness, Jo; Rongved, Pal; Hogset, Anders; Tolleshaug, Helge; Cuthbertson, Alan; et al.

PATENT ASSIGNEE(S):

Marsden, John Christopher, UK; Nycomed Imaging AS

SOURCE:

PCT Int. Appl., 150 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

NT: 10

FAMILY ACC. NUM. COUNT:

PA	PATENT NO.					KIND DA'		DATE A			APPLICATION NO.					ATE		
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WO	9818	500			A2		1998	0507	1	WO 1	997-0	GB29	53		19	9971	028 <	
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		VN,	YU,	ZW														
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CA 2269985					A1	A1 19980507			(CA 1997-2269985					19971028 <			
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PRIORITY APPLN. INFO.:
                                            GB 1996-22366
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                                                                 W 19971028 <--
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                                            US 2001-925715
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AB Targetable diagnostic and/or therapeutically active agents, e.g. ultrasound contrast agents, comprising a suspension in an aqueous carrier liquid of a reporter comprising gas-containing or gas-generated material, in which the reporter is coupled or linked to one or more non-bioactive vectors. Thus, lipopeptide R-Lys(R)-Lys-Arg-Lys-Arg-Trp-Glu-Pro-Pro-Arg-Ala-Arg-Ile- OH (I; R = hexadecanoyl) (preparation given) containing a heparin binding site and a fibronectin binding site, was prepared by standard solid-phase methods. Microbubbles containing lipopeptide I were tested in vitro for binding to endothelial cells under flow conditions.

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L36 ANSWER 18 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 18
ACCESSION NUMBER:
                        1998:300864 HCAPLUS Full-text
```

DOCUMENT NUMBER:

129:4870

TITLE:

Preparation of targetable diagnostic and therapeutic

ultrasound contrast agents

INVENTOR(S):

Klaveness, Jo; Rongved, Pal; Hogset, Anders; Tolleshaug, Helge; Godal, Aslak; Cuthbertson,

Alan; et al.

PATENT ASSIGNEE(S):

Marsdan, John Christopher, UK; Nycomed Imaging AS

SOURCE:

PCT Int. Appl., 140 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT: 10

PATENT INFORMATION:

PA	TENT	NO.			KIN		DATE		į	APPL	ICAT	ION 1	. OI		Ι	ATE		
 MC	9818	100			A2		1998	~ 	,	WO 1	007 /	 cp201	·		-	0071	020	<
WC			7. M	ית ע			BA,							CNT				
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							SE,											
							SE,	SG,	51,	SK,	SL,	10,	1141,	IK,	11,	UA,	UG,	
	DW.				YU,		SZ,	IIC	7 W	א ידי	סס	СП	שת	שת	E.C	. БТ	מש	
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. EF		DE,	D.C	гD		TT	1993	1213		GP 1.	J	J10J.	10		L		026	<
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	6331				B1		2000			по 1. По 1.	007	9592	16			.9971 .9971		
	1442				A1		2001		,			7226				.9971 .9980		
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	к.		FI,		DE,	DIC,	ES,	FR,	GB,	GR,	11,	ш.,	шо,	1411,	SE,	MC,	Ρ1,	
FC	2224	•	гт,	Cı	Т3		2005	0201	,	EC 1	000_	91746	5 1		,	.9980	121	_
	2002		17		A1		2003					9257:				20010		
	6680		1,		B2		2002			U.S Z	001	<i>3431</i> .	LJ		2	.0010	010	~
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PRIORIT					AT		2003	0100				22364				.9961		
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												698				.9970		
												699				.9970		
												8265				.9970		
												11842				.9970		
												11844				.9970		
												49264				9970		
												22369				9961		
												2195			A 1	9970	204	<
									. (GB 1:	997-:	1183	7		A 1	9970	606	<
						`			(GB 1:	997-	11839	9		A 1	9970	606	<
									1	US 1:	997-4	49263	3 P	:	P 1	.9970	607	<
												49266			P 1	9970	607	<
							,					95920			A 1	9971	028 [.]	<
									Ī	WO 1	997-0	GB299	58	1	W 1	9971	028	<
]	EP 1	998-	91746	51		A3 1	9980	424	<
									ι	US 2	001-	9257	15	2	A1 2	0010	810	<
AB T	argeta	able	diag	nost	ic a	nd/	or th	nerap	euti	.call	у ас	tive	age	nts,	е.	g. ul	ltra	sound

contrast agents, comprising a suspension in an aqueous carrier liquid of a reporter comprising gas-containing or gas-generated material, in which the reporter is coupled or linked to one or more non-bioactive vectors. Thus, a mixture of phosphatidylserine, phosphatidylcholine, and biotinamidocaproate-PEG3400-L-Ala-cholesterol (preparation given) was dispersed in 5% propylene glycol-water, flushed with perfluorobutane, and sonicated to give gas-filled encapsulated microbubbles.

L36 ANSWER 19 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 19

ACCESSION NUMBER:

1998:300863 HCAPLUS Full-text

DOCUMENT NUMBER:

129:4869

TITLE:

Preparation of endothelin receptor-binding ultrasound

contrast agents

INVENTOR(S): Klaveness, Jo; Naevestad, Anne; Cuthbertson,

Alan; Solbakken, Magne

PATENT ASSIGNEE(S):

Nycomed Imaging AS, Norway; Cockbain, Julian

SOURCE:

PCT Int. Appl., 98 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

10

PATENT INFORMATION:

PAT	rent n	10.		KIND DATE			APPLICATION NO.						DATE					
WO	98184	97			A2	-		0507					1	9971	028	<		
WO	98184	97			A 3		1998	0716										
	W:	AL,	AM,	AT,	AU,	AZ,	BB,	BG,	BR,	BY	CA,	CH,	CN,	CU,	CZ,	DE,	DK	,
											ıs,							
											MN,							
		RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TR,	TT,	UA,	UG,	US,	UZ,	,
		VN,	YU,	zw														
	RW:	GH,	KE,	LS,	MW,	SD,	SZ,	UG,	ZW,	AT,	BE,	CH,	DE,	DK,	ES,	FI,	FR	,
		GB,	GR,	IE,	IT,	LU,	MC,	NL,	PT,	SE	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	,
		GN,	ML,	MR,	NE,	SN,	TD,	TG										
AU	97478	69			Α		1998	0522	,	AU :	L997-	4786	9		1	9971	028	<
	94620				A2		1999	1006		EP :	L997-	9105	17		1	9971	028	<
EP	94620				В1		2003											
		AT, IE,		CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT	,
US	63312				В1		2001	1218		us -	1997-	95921	0.6		1	9971	028	<i></i>
	24924				T			0915			L997-					9971		
	22066				T3		2004				L997-					9971		
US	20021	022	17		A1		2002				2001-			2001				
US	66800	47			В2		2004	0120										
US	20050	028	65		A1		2005	0106		us 2	2003-	7347	3 0		2	0031	215	<
PRIORITY	Y APPL	N. :	INFO	. :						GB :	L996-:	22364	4			9961		
									(GB :	L996-	2236!	5		A 1	9961	028	<
										GB :	1996-	2236	6		A 1	9961	028	< ~ -
										GB :	L996-	2236	7		A 1	9961	028	<
									(GB :	1996-	2236	В		A 1	9961	028	<
									(GB :	L996-	2236	9	-	A 1	9961	028	<
										GB :	L997-	699		4	A 1	9970	115	<
											L997-					9970		
											L997-					9970		
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											2001-					0010		
OTHER SO	OURCE (s) :			MARI	РΑТ	129.	4869						•		3010	010	•

OTHER SOURCE(S): MARPAT 129:4869

Compns. of matter V-L-R (V is a non-peptidic organic group having binding affinity for an endothelin receptor site; L is a linker moiety or a bond; R is a moiety detectable in in vivo imaging of a human or animal body) are described. Thus, syntheses of Gd(III) and Tc chelates of a DPTA conjugate of a lysine conjugate of 27-0-3-[2-(3-carboxyacryloylamino)-5hydroxyphenyl]acryloyloxymyricerone are described.

L36 ANSWER 20 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 20

ACCESSION NUMBER: 1998:304262 HCAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 129:2225

TITLE: Contrast agents

INVENTOR(S): Klaveness, Jo; Naevestad, Anne; Cuthbertson,

Alan

PATENT ASSIGNEE(S): Nycomed Imaging A/S, Norway; Cockbain, Julian

SOURCE: PCT Int. Appl., 99 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 10

PA	PATENT NO.				KIND DATE			APPLICATION NO.										
WO	EE, LC, RO,	ES, LK,	FI, LR, SD,	A2 AU, GB, LS,	AZ, GE, LT,	1998 BB, GH, LU,	0507 BG, HU, LV,	7 WO 1997-GB2956 , BR, BY, CA, CH, CN, , ID, IL, IS, JP, KE, , MD, MG, MN, MW, MX, , SL, TJ, TM, TR, TT,				56 CN, KE, MX,	CU, KG, NO,	CZ, DE, DK, KP, KR, KZ, NZ, PL, PT,			,	
	GB,	GR,	IE,	IT,	LU,	SZ, MC, TD,	NL,											
AU	9747868	,	,	A		1998			ΙIΑ	19	97-4	1786	8		1	9971	028	<
	971747			A2		2000										9971		
	971747			B1		2005									-	,,,	020	
			CH,			ES,		GB,	GF	R,	IT,	LI,	LU,	NL,	SE,	MC,	PT	,
	IE,	FI																
US	6331289			B1		2001	1218		US	19	997-9	9592	06		1	9971	028	<
AT	314097			${f T}$		2006	0115		AT	19	97-9	9105	16		1	9971	028	<
ES	2257771			T3		2006	0801				997-9					9971	028	<
US	6264914			B1		2001	0724		US	19	99-3	3004	34		1	9990	428	<
US	20010165	87		Al		2001	0823		US	20	01-1	7851	77		2	0010	220	<
US	6524552			B2		2003	0225											
US	20021022	217		A1		2002	0801		US	20	01-9	9257	15		2	0010	810	<
US	6680047			B2		2004	0120											
US	20032282	254		A1		2003	1211		US	20	03-3	3700	92		2	0030	221	<
US	6921525			B2		2005	0726											
US	20050028	365		A1	•	2005	0106		US	20	03-1	7347	30		2	0031	215	<
US	20052019	930		A1		2005	0915				05-3					0050		
	7182934			В2		2007												
PRIORIT	Y APPLN.	INFO	. :						GB	19	96-2	2236	4		A 1	9961	028	<
									GB	19	96-2	2236	5		A 1	9961	028	<
									GB	19	96-2	2236	6			9961		
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	•										97-6					9970		
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											97-6					9970		
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US	1997-959206	Α	19971028	<
WO	1997-GB2956	W	19971028	<
US	1999-300434	А3	19990428	<
US	2001-785177	А3	20010220	<
US	2001-925715	A1	20010810	<
US	2003-370092	A3	20030221	<

OTHER SOURCE(S): MARPAT 129:2225

AB The invention provides a composition of matter (I): V-L-R where V is an organic group having binding affinity for an angiotensin II receptor site, L is a linker moiety or a bond, and R is a moiety detectable in in vivo *imaging* of a human or animal body, with the provisos that where V is angiotensin or a peptidic angiotensin derivative or analog then V-L-R is other than a nonmetal radionuclide substituted peptide (e.g. 125I substituted angiotensin II) and L-V is other than simply a peptide with a chelating agent amide bonded to a side chain thereof. This composition of matter may be used to image cardiovascular diseases and disorders.

L36 ANSWER 21 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2007:173633 HCAPLUS Full-text

TITLE:

Separation processes

INVENTOR(S):

Rongved, Pal; Loevhaug, Dagfinn; Fjerdingstad, Hege;

Solbakken, Magne; Godal, Aslak;

Cuthbertson, Alan

PATENT ASSIGNEE(S):

Norway

SOURCE:

U.S. Pat. Appl. Publ., 17pp., Cont.-in-part of U.S.

Ser. No. 722,075.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 10

PATENT NO.		DATE	APPLICATION NO.	DATE
US 2007036722 CN 1234742 HU 9904595 US 6331289 AT 318618 US 6261537 EP 1442751	A1 A A2 B1 T B1 A1 DE, DK	20070215 19991110 20000428 20011218 20060315 20010717 20040804 , ES, FR, GE	US 2006-498651 CN 1997-199047 HU 1999-4595 US 1997-959206 AT 1997-910514 US 1997-960054	20060803 < 19971028 < 19971028 < 19971028 < 19971028 < 19971029 < 19980424 < SE, MC, PT,
US 2002102215 US 2002102217 US 6680047 CN 1440816 US 2004141922 US 2005002865 PRIORITY APPLN. INFO.:	A1 B2 A A1	20020801 20020801 20040120 20030910 20040722 20050106	US 2001-765614 US 2001-925715 CN 2002-160420 US 2003-722075 US 2003-734730 GB 1996-22366 GB 1996-22367 GB 1996-22368 GB 1997-699 GB 1997-8265 GB 1997-11842 GB 1997-11846 US 1997-49264P	20010810 < 20021230 < 20031126 <

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US 1997-49265P
                    P 19970606 <--
US 1997-49268P
                    P 19970606 <--
US 1997-958993
                    A2 19971028 <--
US 1997-960054
                    A1 19971029 <--
US 2001-765614
                    B1 20010122 <--
US 2003-722075
                    A2 20031126 <--
GB 1996-22369
                    A 19961028 <--
GB 1997-2195
                    A 19970204 <--
GB 1997-11837
                    A 19970606 <--
GB 1997-11839
                    A 19970606 <--
US 1997-49263P
                    P
                       19970607 <--
US 1997-49266P
                    P 19970607 <--
US 1997-959206
                    A 19971028 <--
EP 1998-917461
                    A3 19980424 <--
US 2001-925715
                    A1 20010810 <--
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AB Separation of target material from a liquid sample is achieved by coupling the target to targetable encapsulated gas microbubbles, allowing the microbubbles and coupled target to float to the surface of the sample to form a floating microbubble/target layer, and separating this layer from the sample. In a pos. separation process the microbubbles are then removed from the target, e.g. by bursting. In a neg. separation process target-free sample material is recovered following separation of the floating layer. The method may also be used diagnostically to detect the presence of a disease marker in a sample. Novel separation apparatus is also described. Gas microbubbles encapsulated with DSPS and thiolated anti-CD34 antibodies-Mal-PEG2000DSPE, useful for separation of hematopoietic stem cells, were prepared

L36 ANSWER 22 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2004:780565 HCAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 141:277892

TITLE: Methods of radiofluorination of peptides and other

biologically active vectors

INVENTOR(S): Cuthbertson, Alan; Solbakken, Magne

; Arukwe, Joseph Maduabuchi; Karlsen, Hege; Glaser,

Matthias Eberhard

PATENT ASSIGNEE(S): Amersham Health AS, Norway; Hammersmith Imanet Ltd.

SOURCE: PCT Int. Appl., 56 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT NO.				KIND DATE		APPLICATION NO.					DATE					
WO 2004080492				A1 20040923			WO 2004-GB1052					20040312 <				
W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	ΒZ,	CA,	CH,
	CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
	GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KΖ,	LC;
	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,
	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,
	ТJ,	TM,	TN,	TR,	TT,	TZ,	ŲΑ,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW
RW:	BW,	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ŻW,	AM,	AZ,
,	BY,	KG,	KZ,	MD,	RU,	ТJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,
	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,	SI,
	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,
	TD,	TG														
AU 2004	2188	79		A1 20040923			AU 2004-218879					20040312 <				
CA 2518889			A1	A1 20040923			CA 2004-2518889					20040312 <				

EP 1601384 A1 20051207 EP 2004-720084 20040312 <--R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK BR 2004008236 Α 20060301 BR 2004-8236 20040312 <--CN 1758925 Α 20060412 CN 2004-80006725 20040312 <--JP 2006523658 Т 20061019 JP 2006-505951 20040312 <--NO 2005004185 Α 20051110 NO 2005-4185 20050909 <--PRIORITY APPLN. INFO.: GB 2003-5704 20030313 <--WO 2004-GB1052 20040312

OTHER SOURCE(S):

MARPAT 141:277892

GT

The invention relates to diagnostic and radic biol. active vectors labeled with positron-emitting nuclides. It rurner relates to methods and reagents for (18F)-fluorination of vectors, where a vector is defined as a mol. with an affinity for a specific biol. target, and is preferably a peptide. The resultant 18F-labeled conjugates are useful as radiopharmaceuticals, specifically for use in Positron Emission Tomog. (PET). Compds. 18F-linker-X-N:C(Y)-vector and vector-X-N:C(Y)-linker-18F (X is CONH, NH, O, NHCONH or NHCSNH; Y is H, alkyl or aryl) are claimed. Thus, I (Boc = tert-butoxycarbonyl), prepared by standard peptide synthesis and coupling with Boc-NHOCH2CO2H, was deprotected and conjugated with 4-18FC6H4CHO.

RETABLE

Referenced Author	Year	VOT	PG	Referenced Work	Referenced
(RAU)	(RPY)	(RVL)	(RPG)	(RWK)	File
=======================================	+=====	+=====	+=====	+====================================	+========
Griffiths, G	1999	1		WO 9911590 A	HCAPLUS
Hwang, D	1991	32	1730	JOURNAL OF NUCLEAR M	HCAPLUS

L36 ANSWER 23 OF 25

HCAPLUS COPYRIGHT 2007 ACS on STN 2003:777733 HCAPLUS Full-text

ACCESSION NUMBER: DOCUMENT NUMBER:

139:277170

TITLE:

Methods and reagents for radiofluorination,

particularly of peptides

INVENTOR (S):

Cuthbertson, Alan; Solbakken, Magne

; Arukwe, Joseph Maduabuchi; Karlsen, Hege

PATENT ASSIGNEE(S):

Amersham PLC, UK

SOURCE:

PCT Int. Appl., 38 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.	
WO 2003080544	A1 20031002	WO 2003-GB1332	
		BA, BB, BG, BR, BY,	
CO, CR, CU,	CZ, DE, DK, DM,	DZ, EC, EE, ES, FI,	GB, GD, GE, GH,
GM, HR, HU,	ID, IL, IN, IS,	JP, KE, KG, KP, KR,	KZ, LC, LK, LR,
LS, LT, LU,	LV, MA, MD, MG,	MK, MN, MW, MX, MZ,	NI, NO, NZ, OM,
PH, PL, PT,	RO, RU, SC, SD,	SE, SG, SK, SL, TJ,	TM, TN, TR, TT,
TZ, UA, UG,	US, UZ, VC, VN,	YU, ZA, ZM, ZW	
RW: GH, GM, KE,	LS, MW, MZ, SD,	SL, SZ, TZ, UG, ZM,	ZW, AM, AZ, BY,
KG, KZ, MD,	RU, TJ, TM, AT,	BE, BG, CH, CY, CZ,	DE, DK, EE, ES,
FI, FR, GB,	GR, HU, IE, IT,	LU, MC, NL, PT, RO,	SE, SI, SK, TR,
BF, BJ, CF,	CG, CI, CM, GA,	GN, GQ, GW, ML, MR;	NE, SN, TD, TG
AU 2003214446	A1 20031008	AU 2003-214446	20030320 <
EP 1487762 '	A1 20041222	EP 2003-710021	20030320 <
EP 1487762	B1 . 20070228		
R: AT, BE, CH,	DE, DK, ES, FR,	GB, GR, IT, LI, LU,	NL, SE, MC, PT,
IE, SI, LT,	LV, FI, RO, MK,	CY, AL, TR, BG, CZ,	EE, HU, SK
US 2005142061	A1 20050630	US 2003-508682	20030320 <
JP 2005520857	T 20050714	JP 2003-578305	20030320 <
PRIORITY APPLN. INFO.:		GB 2002-6750	A 20020322 <
		WO 2003-GB1332	W 20030320 <
OTHER SOURCE(S):	MARPAT 139:2771	70	
GI			

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB The invention relates to the synthesis of 18F-labeled compds., particularly peptides, for use as radiopharmaceuticals, specifically for use in positron emission tomog. (PET). The radiofluorination method involves reacting a compound X-CH2CONH-peptide (X is a halogen leaving group, preferably chloro) or maleimido-Y-CONH-peptide (Y is a C1-10 hydrocarbyl group optionally containing 1-6 heteroatoms) with a compound 18F-(linker)-SH, in which the linker is a C1-30 hydrocarbyl group optionally containing 1-10 heteroatoms. Thus, compound I was prepared by site-specific conjugation of 4-FCH2C6H4CONHCH2CH2SCPh3 to the maleimide-modified peptide.

RETABLE

Referenced Author (RAU)	(RPY) (PG (RPG)	Referenced Work (RWK)	Referenced File
Dean, R Griffiths, G Hwang, D	1992 1999	 		US 5144043 A	HCAPLUS HCAPLUS

L36 ANSWER 24 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 1999:708880 HCAPLUS Full-text

DOCUMENT NUMBER:

131:319884

TITLE: Targetable encapsulated gas microbubbles for

separation of target material from liquid samples and

separation apparatus

INVENTOR(S): Cuthbertson, Alan; Rongved, Pal; Lovhaug,

Dagfinn; Fjerdingstad, Hege; Solbakken, Magne

; Godal, Aslak

PATENT ASSIGNEE(S): Nycomed Imaging As, Norway

SOURCE:

PCT Int. Appl., 54 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	TENT										LICAT							•
WO	9955	837			A2		1999	1104			 1999-					9990		
WO	9955	837			A3		2000	0210										
	W :	ΑE,	AL,	AM,	ΑT,	ΑU,	ΑZ,	ΒA,	BB,	BG	, BR,	BY,	CA,	CH,	CN,	CU,	CZ	
		DE,	DK,	EE,	ES,	FI,	GB,	GD,	GE,	GH	, GM,	HR,	HU,	ID,	IL,	IN,	IS,	
		JP,	KE,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR	, LS,	LT,	LU,	LV,	MD,	MG,	MK,	
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	RW:	GH,	GM,	KΕ,	LS,	MW,	SD,	SL,	SZ,	UG	, ZW,	ΑT,	BE,	CH,	CY,	DE,	DK,	
		ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC	, NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	
		CI,	CM,	GA,	GN,	GW,	ML,	MR,	NE,	sn	, TD,	TG		•				
CA	2326	386			A1		1999	1104		CA	1999-	2326	386		1	9990	428	<
AU	9937	197			Α		1999	1116		ΑU	1999-	3719	7		1	9990	428	<
EP	1073	716			A2		2001	0207		ΕP	1999-	9193	96		1	9990	428	<
EP	1073	716			B1		2004	0428										
	R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR	, IT,	LI,	LU,	NL,	SE,	MC,	PT,	
		ΙĒ,	FI															
JP	2002	5128	86		T		2002	0508		JP .	2000-	5459	81		1	9990	428	<
AΤ	2655	25			T		2004	0515		ΑT	1999-	9193	96		1	9990	428	<
IN	2000	00MM	515		Α		2005	0715		IN.	2000-	MN51	5		2	0001	018	<
NO	2000	0053	83		Α	:	2000	1213		NO :	2000-	5383			2	0001	026	<
US	2003	1043	59		A1		2003	0605	•	US :	2002-	2945	98		2	0021	115	<
IORIT	Y APP	LN.	INFO	.:						GΒ	1998-	9083		7	A 1	9980	428	<
									4	GB	1998-	9085		7	A 1	9980	428	<
									•	US.	1998-	8581	9P]	P 1	9980	518	<
										US	1998-	8582	6P]	P 1	9980	518	<
									1	WO	1999-	GB13	17	1	W 1	9990	428	<
											2000-					•		
Se	para	tion	of t	arge	t ma	teri	lal f	rom	a li	.gui	d san	nple	is a	chie	ved	by c	guo	lino

AB Separation of target material from a liquid sample is achieved by coupling the target to targetable encapsulated gas microbubbles, allowing the microbubbles and coupled target to float to the surface of the sample to form a floating microbubble/target layer, and separating this layer from the sample. In a pos. separation process the microbubbles are then removed from the target, e.g. by bursting. In a neg. separation process target-free sample material is recovered following separation of the floating layer. The method may also be used diagnostically to detect the presence of a disease marker in a sample. Novel separation apparatus is also described. Perfluorobutane gas microbubbles encapsulated with distearoylphosphatidylserine doped with Mal-PEG2000-distearoylphosphatidylethanolamine (DSPE) was prepared and reacted with thiolated anti-CD34 antibodies to make a reagent useful for separating CD34-pos. cells.

L36 ANSWER 25 OF 25 WPIX COPYRIGHT 2007 THE THOMSON CORP on STN ACCESSION NUMBER: 2004-625405 [60] WPIX

DOC. NO. CPI:

C2004-224919 [60]

TITLE:

Diagnostic imaging agent, useful for

diagnostic imaging of cardiovascular and inflammatory diseases, comprises a matrix metalloproteinase inhibitor

labeled with a gamma-emitting radionuclide

DERWENT CLASS:

B04; B05; K08

INVENTOR:

ARUKWE J; BREYHOLZ H; CUTHBERTSON A; DAVIS J;

HEYWOOD K; KOPKA K; LEVKAU B; MENDIZABAL M; RICKETTS S; SCHAFERS M; STOREY A; STOREY A E; WAGNER S; WILSON I;

WYNN D

PATENT ASSIGNEE:

(AMSH-C) AMERSHAM PLC; (GENE-C) GE HEALTHCARE LTD

COUNTRY COUNT:

107

PATENT INFO ABBR.:

PAT	TENT NO	KINI	DATE	WEEK	LA	PG	MAIN	IPC	
÷									
WO	2004069365	A1	20040819	(200460)*	EN	61[12]		•	
ΑU	2004210208	A1	20040819	(200559)	EN				
ΕP	1592458	A1	20051109	(200573)	EN				
NO	2005003776	Α	20050930	(200574)	NO				
CN	1747749	Α	20060315	(200649)	ZH				
JP	2006519216	W	20060824	(200656)	JA	57			

APPLICATION DETAILS:

PATENT NO KIND	APPLICATION DATE
WO 2004069365 A1	WO 2004-GB524 20040210
AU 2004210208 A1	AU 2004-210208 20040210
CN 1747749 A	CN 2004-80003837 20040210
EP 1592458 A1	EP 2004-709657 20040210
EP 1592458 A1	WO 2004-GB524 20040210
NO 2005003776 A	NO 2005-3776 20050809
JP 2006519216 W	WO 2004-GB524 20040210
JP 2006519216 W	JP 2006-502261 20040210

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 2004210208	A1 Based	on WO 2004069365 A
EP 1592458	A1 Based	on WO 2004069365 A
JP 2006519216	W Based	on WO 2004069365 A

PRIORITY APPLN. INFO: GB 2003-7524 20030401
GB 2003-2891 20030210

AN 2004-625405 [60] WPIX

AB WO 2004069365 A1 UPAB: 20060203

NOVELTY - Diagnostic *imaging agent* (A) comprises a matrix metalloproteinase inhibitor (I) labeled with a gamma-emitting radionuclide.

DETAILED DESCRIPTION - Diagnostic *imaging agent* (A) comprises a matrix metalloproteinase inhibitor of formula (I) labeled with a gamma-emitting radionuclide.

Either R1 = H, OH, 1-6C alkyl, 6-14C aryl or 7-20C arylalkyl; or C+R1+R5 = 6-8C cycloalkyl ring or a 4-6C heterocyclic ring; or

C+R1+R4=4-6C heterocyclic ring containing 5-7 atoms and 1 or 2 heteroatoms from N or O;

R2, R3 = H, OH, halo, 1-6C alkyl, 1-6C alkoxy, 1-6C amino, 6-14C aryl, $^{\circ}$ 7-20C arylalkyl or 7-20C carbamoylaryl;

R4 = 6-14C aryl, 4-6C heteroaryl, 7-20C arylalkyl, 7-20C carbamoylaryl or arylcarbamoylaryl; and

R5 = H or 1-6C alkyl.

INDEPENDENT CLAIMS are also included for:

- (1) a ligand conjugate which comprises (I), conjugated to a ligand suitable for the co-ordination of a gamma-emitting radio metal (99m-Tc, 111ln, 113mln, 67Cu or 67Ga);
- (2) a precursor useful in the preparation of (A) comprising a group suitable for reaction with a gamma-emitting isotope of iodine to give (A);
- (3) a pharmaceutical composition comprising (A) with a biocompatible carrier; and
- (4) a kit for the preparation of a pharmaceutical composition of (A). ACTIVITY - Cardiovascular-Gen.; Antiarteriosclerotic; Antiinflammatory; Respiratory-Gen.

MECHANISM OF ACTION - Matrix metalloproteinase (MMP) inhibitor.

Compounds (I) were tested for their MMP inhibitory activity using biological assays. The result showed that the median inhibitory concentration value of (2R)-N-hydroxy-2-(((4-iodophenyl)sulfonyl)(pyridin- 3-ylmethyl)amino)-3-methylbutanamide was 2.5 nM.

USE - (A) is useful for the diagnostic imagining of cardiovascular disease (preferably atherosclerosis and congestive heart failure) and inflammatory diseases (preferably chronic obstructive pulmonary disease) (claimed).

=> d que 19

L1 STR

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

Structure attributes must be viewed using STN Express query preparation.

L3 1 SEA FILE=HCAPLUS ABB=ON PLU=ON US2006-566487/AP

L5 4 SEA FILE=REGISTRY SSS FUL L1

L8 1 SEA FILE=HCAPLUS ABB=ON PLU=ON L5

L9 1 SEA FILE=HCAPLUS ABB=ON PLU=ON (L8 OR L3)

=> d ibib abs hitstr retable 19 tot

L9 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:120957 HCAPLUS Full-text

DOCUMENT NUMBER: 142:219561

TITLE: Preparation of peptide-based compounds as diagnostic

imaging agents

INVENTOR(S): Cuthbertson, Alan; Solbakken, Magne

PATENT ASSIGNEE(S): Amersham Health AS, Norway

SOURCE: PCT Int. Appl., 40 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005012335	A1	20050210	WO 2004-GB3150	20040721
W: AE, AG,	AL, AM, AT	r, Au, Az,	BA, BB, BG, BR, BW, BY,	BZ, CA, CH,
CN, CO,	CR, CU, CZ	Z, DE, DK,	DM, DZ, EC, EE, EG, ES,	FI, GB, GD,

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GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
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             SN, TD, TG
    CA 2533321
                          A1
                                20050210
                                             CA 2004-2533321
                                                                     20040721
    EP 1648925
                                20060426
                                             EP 2004-743485
                          A1
                                                                    20040721
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK
    CN 1829735
                          Α
                                20060906
                                             CN 2004-80022044
                                                                    20040721
    BR 2004012986
                          Α
                                20061003
                                             BR 2004-12986
                                                                    20040721
    HU 200600230
                          A2
                                20070129
                                             HU 2006-230
                                                                    20040721
    US 2006193773
                          Α1
                                20060831
                                             US 2006-566487
                                                                    20060130 <--
    NO 2006000825
                                20060329
PRIORITY APPLN. INFO.:
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OTHER SOURCE(S):

CASREACT 142:21956

GΤ

Ι

The invention relates to compds. I [R2 is [NH(CH2CH2O)3CH2CH2NHCOCH2CO]0-10NH2; R3 is an alkylene or alkenylene bridge; W1 is absent or a spacer moiety (hetero)hydrocarbyl preferably derived from glutaric and/or succinic acid and/or a polyethylene glycol-based unit and/or a unit [NH(CH2CH2O)3CH2CH2NHCOCH2CO]n; Z1 is an antineoplastic agent, a chelating agent or a reporter moiety] and their use as targeting vectors that bind to receptors associated with angiogenesis. Compds. I may thus be used for diagnosis or therapy of various diseases. Thus, compound I [R2 is NH2, R3 is CH2, Z1-W1 is FCH2CH2SCH2CONH(CH2CH2O)5CH2CH2NHCOCH2COH2CONH] was prepared via the solid-phase method and showed Ki = 7 nmol in an $\alpha\nu\beta3$ integrin receptor binding assay.

IT 840474-71-7P 840474-72-8P

RL: DGN (Diagnostic use); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of peptide-based compds. as diagnostic imaging agents)

RN 840474-71-7 HCAPLUS

CN L-Cysteinamide, N6-(30-fluoro-1,5,25-trioxo-3,9,12,15,18,21-hexaoxa-27-thia-6,24-diazatriacont-1-yl)-N2-(mercaptoacetyl)-L-lysyl-S (mercaptomethyl)-L-cysteinyl-L-arginylglycyl-L-α-aspartyl-L cysteinyl-L-phenylalanyl-, cyclic (1→8),(2→6)-bis(thioether)
 (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 840474-72-8 HCAPLUS

CN L-Cysteinamide, N6-[30-(fluoro-18F)-1,5,25-trioxo-3,9,12,15,18,21-hexaoxa27-thia-6,24-diazatriacont-1-yl]-N2-(mercaptoacetyl)-L-lysyl-S(mercaptomethyl)-L-cysteinyl-L-arginylglycyl-L-α-aspartyl-Lcysteinyl-L-phenylalanyl-, cyclic (1→8),(2→6)-bis(thioether)
(9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

IT 840474-69-3P 840474-70-6P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of peptide-based compds. as diagnostic imaging agents)

RN 840474-69-3 HCAPLUS

CN L-Cysteinamide, N2-(mercaptoacetyl)-L-lysyl-S-(mercaptomethyl)-L-cysteinyl-L-arginylglycyl-L- α -aspartyl-L-cysteinyl-L-phenylalanyl-, cyclic (1 \rightarrow 8), (2 \rightarrow 6)-bis(thioether) (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 840474-70-6 HCAPLUS

CN L-Cysteinamide, N6-(26-chloro-1,5,25-trioxo-3,9,12,15,18,21-hexaoxa-6,24-diazahexacos-1-yl)-N2-(mercaptoacetyl)-L-lysyl-S-(mercaptomethyl)-L-cysteinyl-L-arginylglycyl-L- α -aspartyl-L-cysteinyl-L-phenylalanyl-, cyclic (1 \rightarrow 8), (2 \rightarrow 6)-bis(thioether) (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE *** RETABLE

Referenced Author (RAU)	Year (RPY)	VOL (RVL)	1 '	Referenced Work (RWK)	Referenced File
		+=====	+=====-	+=============	+========
Bonasera, T	2002	}		WO 02062819 A	HCAPLUS
Harris, T	1996	6 .	1741	BIOORGANIC & MEDICIN	HCAPLUS
Indrevoll, B	2001	ļ [*]	 -	WO 0177145 A	HCAPLUS
Indrevoll, B	2003	1	1	WO 03006491 A	HCAPLUS
Lister-James, J	1999		İ	US 5888474 A	HCAPLUS
Pearson, D	1996	39	1372	JOURNAL OF MEDICINAL	HCAPLUS
Srinivasan, A	2002	Ι .	1	WO 0220610 A	HCAPLUS

=> d his full

L1

(FILE 'HOME' ENTERED AT 13:43:12 ON 15 MAR 2007)

FILE 'REGISTRY' ENTERED AT 13:43:18 ON 15 MAR 2007 STRUCTURE UPLOADED

L2 0 SEA SSS SAM L1

FILE 'HCAPLUS' ENTERED AT 13:43:41 ON 15 MAR 2007

E US2006-566487/APPS

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L3
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                D SCAN
                SEL RN L3
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L4
                541-88-8/BI OR 606975-95-5/BI OR 606975-96-6/BI OR 606976-02-7/
                BI OR 840474-68-2/BI OR 840474-69-3/BI OR 840474-70-6/BI OR
                840474-71-7/BI OR 840474-72-8/BI)
                D SCAN
                D QUE L1
L5
              4 SEA SSS FUL L1
                D SCAN
                SAVE L5 YOUNG487/A TEMP
              4 SEA ABB=ON PLU=ON L4 AND L5
                D SCAN
T.7
              7 SEA ABB=ON PLU=ON L4 NOT L6
                D SCAN
     FILE 'HCAPLUS' ENTERED AT 13:45:59 ON 15 MAR 2007
L8
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L9
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     FILE 'BEILSTEIN' ENTERED AT 13:46:21 ON 15 MAR 2007
L10
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     FILE 'MARPAT' ENTERED AT 13:46:33 ON 15 MAR 2007
L11
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              2 SEA SSS FUL L1
L12
L13
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              O SEA ABB=ON PLU=ON L13 NOT L9
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L15
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                "CUTHBERTSON ALAN J S"/AU OR "CUTHBERTSON ALAN S"/AU OR
                "CUTHBERTSON ALLAN S"/AU)
                E SOLBAKKEN M/AU
L16
             28 SEA ABB=ON PLU=ON ("SOLBAKKEN M"/AU OR "SOLBAKKEN MAGNE"/AU)
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L17
L18
           14 SEA ABB=ON PLU=ON L17 AND (AY<2004 OR PY<2004 OR PRY<2004)
L19
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L20
             24 SEA ABB=ON PLU=ON (L20 OR L18)
L21
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    ON 15 MAR 2007
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L23
             74 SEA ABB=ON PLU=ON SOLBAKKEN M?/AU
            49 SEA ABB=ON PLU=ON L22 AND L23
L24
            36 SEA ABB=ON PLU=ON L24 AND IMAGING?
25 SEA ABB=ON PLU=ON L25 AND (AY<2004 OR PY<2004 OR PRY<2004)
L25
L26
L27
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L28
            29 SEA ABB=ON PLU=ON L27 AND (AY<2004 OR PY<2004 OR PRY<2004)
L29
            40 SEA ABB=ON PLU=ON (L26 OR L28)
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L30 29 SEA ABB=ON PLU=ON L29 AND (IMAGING AGENT?)

FILE 'STNGUIDE' ENTERED AT 13:53:33 ON 15 MAR 2007

FILE 'REGISTRY' ENTERED AT 13:55:15 ON 15 MAR 2007

D RSD L6 TOT

L31 4 SEA ABB=ON PLU=ON 105465.1/RID

L32 4 SEA ABB=ON PLU=ON L31 AND 46.150/RID

L33 4 SEA ABB=ON PLU=ON (L31 OR L32)

L34 4 SEA ABB=ON PLU=ON (L6 OR L33)

FILE 'MEDLINE, EMBASE, BIOSIS, DRUGU, CAOLD' ENTERED AT 13:57:13 ON 15 MAR 2007

L35 0 SEA ABB=ON PLU=ON L5 .

FILE 'STNGUIDE' ENTERED AT 13:57:23 ON 15 MAR 2007

D QUE L21

D QUE L30

FILE 'HCAPLUS, MEDLINE, EMBASE, BIOSIS, WPIX' ENTERED AT 13:57:33 ON 15 MAR 2007

L36 25 DUP REM L21 L30 (28 DUPLICATES REMOVED)

ANSWERS '1-24' FROM FILE HCAPLUS

ANSWER '25' FROM FILE WPIX

D IBIB ABS HITSTR RETABLE L36 TOT

D QUE L9

D IBIB ABS HITSTR RETABLE L9 TOT

FILE HOME

FILE REGISTRY

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 14 MAR 2007 HIGHEST RN 926494-79-3 DICTIONARY FILE UPDATES: 14 MAR 2007 HIGHEST RN 926494-79-3

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH December 2, 2006

Please note that search-term pricing does apply when conducting ${\tt SmartSELECT}$ searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

FILE HCAPLUS

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FILE COVERS 1907 - 15 Mar 2007 VOL 146 ISS 12 FILE LAST UPDATED: 14 Mar 2007 (20070314/ED)

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

FILE BEILSTEIN
FILE LAST UPDATED ON JANUARY 10, 2007

FILE COVERS 1771 TO 2006.

FILE CONTAINS 9,780,003 SUBSTANCES

>>>PLEASE NOTE: Reaction Data and substance data are stored in separate documents and can not be searched together in one query. Reaction data for BEILSTEIN compounds may be displayed immediately with the display codes PRE (preparations) and REA (reactions). A substance answer set retrieved after the search for a chemical name, a compounds with available reaction information by combining with PRE/FA, REA/FA or more generally with RX/FA. The BEILSTEIN Registry Number (BRN) is the link between a BEILSTEIN compound and belonging reactions. For mo detailed reaction searches BRNs can be searched as reaction partner BRNs Reactant BRN (RX.RBRN) or Product BRN (RX.PBRN).<<<

>>> FOR SEARCHING PREPARATIONS SEE HELP PRE <<<

- * PLEASE NOTE THAT THERE ARE NO FORMATS FREE OF COST.
- * SET NOTICE FEATURE: THE COST ESTIMATES CALCULATED FOR SET NOTICE
- * ARE BASED ON THE HIGHEST PRICE CATEGORY. THEREFORE; THESE
- * ESTIMATES MAY NOT REFLECT THE ACTUAL COSTS.
- * FOR PRICE INFORMATION SEE HELP COST

- * PATENT NUMBERS (PN) AND BABS ACCESSION NUMBERS (BABSAN) CAN NOW BE SEARCHED, SELECTED AND TRANSFERRED.
- * NEW DISPLAY FORMATS ALLREF, ALLP AND BABSAN SHOW ALL REFERENCES, ALL PATENT REFERENCES, OR ALL BABS ACCESSION NUMBERS FOR A COMPOUND AT A GLANCE.

FILE MARPAT

FILE CONTENT: 1961-PRESENT VOL 146 ISS 11 (20070309/ED)

SOME MARPAT RECORDS ARE DERIVED FROM INPI DATA FOR 1961-1987

MOST RECENT CITATIONS FOR PATENTS FROM MAJOR ISSUING AGENCIES (COVERAGE TO THESE DATES IS NOT COMPLETE):

US 2007020715 25 JAN 2007
DE 102005032918 18 JAN 2007
EP 1743897 17 JAN 2007
JP 2007016265 25 JAN 2007
WO 2007012422 01 FEB 2007
GB 2427406 27 DEC 2006

FR 2888248 12 JAN 2007 RU 2291880 20 JAN 2007 CA 2551930 08 JAN 2007

Expanded G-group definition display now available.

FILE MEDLINE

FILE LAST UPDATED: 14 Mar 2007 (20070314/UP). FILE COVERS 1950 TO DATE.

All regular MEDLINE updates from November 15 to December 16 have been added to MEDLINE, along with 2007 Medical Subject Headings (MeSH(R)) and 2007 tree numbers.

The annual reload will be available in early 2007.

This file contains CAS Registry Numbers for easy and accurate substance identification.

FILE EMBASE

FILE COVERS 1974 TO 14 Mar 2007 (20070314/ED)

EMBASE is now updated daily. SDI frequency remains weekly (default) and biweekly.

This file contains CAS Registry Numbers for easy and accurate substance identification.

FILE BIOSIS

FILE COVERS 1969 TO DATE.

CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 14 March 2007 (20070314/ED)

FILE DRUGU

FILE LAST UPDATED: 15 MAR 2007 <20070315/UP>

>>> DERWENT DRUG FILE (SUBSCRIBER) <<<

>>> FILE COVERS 1983 TO DATE <<<

>>> THESAURUS AVAILABLE IN /CT <<<

FILE WPIX

FILE LAST UPDATED: 14 MAR 2007 <20070314/UP>
MOST RECENT THOMSON SCIENTIFIC UPDATE: 200718 <200718/DW>

DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

>>> New reloaded DWPI Learn File (LWPI) available as well <<<

>>> YOU ARE IN THE NEW AND ENHANCED DERWENT WORLD PATENTS INDEX <<<

>>> New display format FRAGHITSTR available <<< SEE ONLINE NEWS and

http://www.stn-international.de/archive/stn_online_news/fraghitstr_ex.pdf

>>> IPC Reform reclassification data for the backfile is being
loaded into the database during January 2007.
There will not be any update date (UP) written for the reclassified
documents, but they can be identified by 20060101/UPIC. <<<</pre>

FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE, PLEASE VISIT:

http://www.stn-international.de/training_center/patents/stn_guide.pdf

FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE http://scientific.thomson.com/support/patents/coverage/latestupdates/

PLEASE BE AWARE OF THE NEW IPC REFORM IN 2006, SEE

http://www.stn-international.de/stndatabases/details/ipc_reform.html and
http://scientific.thomson.com/media/scpdf/ipcrdwpi.pdf

>>> FOR DETAILS ON THE NEW AND ENHANCED DERWENT WORLD PATENTS INDEX PLEASE SEE

http://www.stn-international.de/stndatabases/details/dwpi r.html <<<

FILE STNGUIDE

FILE CONTAINS CURRENT INFORMATION.

LAST RELOADED: Mar 9, 2007 (20070309/UP).

FILE CAOLD

FILE COVERS 1907-1966

FILE LAST UPDATED: 01 May 1997 (19970501/UP)

This file contains CAS Registry Numbers for easy and accurate substance identification. Title keywords, authors, patent assignees, and patent information, e.g., patent numbers, are now searchable from 1907-1966. TIFF images of CA abstracts printed between 1907-1966 are available in the PAGE display formats.

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file supports REG1stRY for direct browsing and searching of all substance data from the REGISTRY file. Enter HELP FIRST for more information.